=> fil reg

FILE 'REGISTRY' ENTERED AT 09:53:35 ON 11 MAR 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 10 MAR 2003 HIGHEST RN 497818-02-7 DICTIONARY FILE UPDATES: 10 MAR 2003 HIGHEST RN 497818-02-7

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

## => d ide can 12

L2 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS

RN **9072-19-9** REGISTRY

CN Fucoidan (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Fucoidin

CN Fucoidine

CN Nemacystus mucilage

MF Unspecified

CI PMS, COM, MAN

PCT Manual registration

LC STN Files: AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, CHEMCATS, CIN, CSCHEM, EMBASE, IFICDB, IFIPAT, IFIUDB, MEDLINE, NAPRALERT, PROMT, TOXCENTER, USPAT2, USPATFULL

## \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

683 REFERENCES IN FILE CA (1962 TO DATE)

39 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA 684 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 138:136255

REFERENCE 2: 138:117807

REFERENCE 3: 138:112440

REFERENCE 4: 138:69631

REFERENCE 5: 138:54454

REFERENCE 6: 138:44563

REFERENCE 7: 138:37388

REFERENCE 8: 138:20954

REFERENCE 9: 138:13533

Jan Delavai Reference Librarian Biotechnology & Chemical Library C№1 1E07 – 703-308-44.98 jan.delaval@uspto.gov REFERENCE 10: 138:8269

```
=> d ide can 13 tot
```

L3 ANSWER 1 OF 5 REGISTRY COPYRIGHT 2003 ACS

RN 213832-60-1 REGISTRY

CN Fucoidan, acetate (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Acetyl fucoidan

MF C2 H4 O2 . x Unspecified

SR CA

LC STN Files: BIOSIS, CA, CAPLUS

CM 1

CRN 9072-19-9

CMF Unspecified

CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 64-19-7 CMF C2 H4 O2

О || но-с-снз

- 3 REFERENCES IN FILE CA (1962 TO DATE)
- 3 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 134:68485

REFERENCE 2: 130:97113

REFERENCE 3: 129:291355

L3 ANSWER 2 OF 5 REGISTRY COPYRIGHT 2003 ACS

RN 158853-89-5 REGISTRY

CN Fucoidan, 3-amino-2-hydroxypropyl ether (9CI) (CA INDEX NAME)

MF C3 H9 N O2 . x Unspecified

PCT Manual registration

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER

CM 1 .

CRN 9072-19-9 CMF Unspecified CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 616-30-8 CMF C3 H9 N O2

```
\begin{array}{c} \text{OH} \\ | \\ \text{HO-CH}_2\text{-CH-CH}_2\text{-NH}_2 \end{array}
```

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 121:271676

L3 ANSWER 3 OF 5 REGISTRY COPYRIGHT 2003 ACS

RN 120574-85-8 REGISTRY

CN Cytidine, 2',3'-dideoxy-, mixt. with fucoidan (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Fucoidan, mixt. contg. (9CI)

FS STEREOSEARCH

MF  $\,$  C9 H13 N3 O3  $\,$  Unspecified

CI MXS

SR CA

LC STN Files: CA, CAPLUS, DRUGPAT, DRUGUPDATES, TOXCENTER

CM 1

CRN 9072-19-9

CMF Unspecified

CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 7481-89-2

CMF C9 H13 N3 O3

Absolute stereochemistry. Rotation (+).

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 110:219081

L3 ANSWER 4 OF 5 REGISTRY COPYRIGHT 2003 ACS

RN 120485-65-6 REGISTRY

CN Fucoidan, mixt. with 2-amino-1,9-dihydro-9-[(2-hydroxyethoxy)methyl]-6H-purin-6-one (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 6H-Purin-6-one, 2-amino-1,9-dihydro-9-[(2-hydroxyethoxy)methyl]-, mixt. contg. (9CI)

MF C8 H11 N5 O3 . Unspecified

CI MXS

SR CA

LC STN Files: CA, CAPLUS, DRUGPAT, TOXCENTER

CM 1

CRN 59277-89-3 CMF C8 H11 N5 O3

$$H_2N$$
 $N$ 
 $H_2N$ 
 $N$ 
 $H$ 
 $CH_2-O-CH_2-CH_2-OH$ 

CM 2

CRN 9072-19-9 CMF Unspecified CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

1 REFERENCES IN FILE CA (1962 TO DATE)
1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 110:219081

L3 ANSWER 5 OF 5 REGISTRY COPYRIGHT 2003 ACS

RN 120485-60-1 REGISTRY

CN Thymidine, 3'-azido-3'-deoxy-, mixt. with fucoidan (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Fucoidan, mixt. contg. (9CI)

FS STEREOSEARCH

MF C10 H13 N5 O4 . Unspecified

CI MXS

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER

CM 1

CRN 30516-87-1 CMF C10 H13 N5 O4

Absolute stereochemistry. Rotation (+).

CM 2

CRN 9072-19-9

CMF Unspecified CCI PMS, MAN

L27

9782 S E38,E39 E E3+ALL

```
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
               1 REFERENCES IN FILE CA (1962 TO DATE)
               1 REFERENCES IN FILE CAPLUS (1962 TO DATE)
          1: 110:219081
REFERENCE
=> d his
     (FILE 'HOME' ENTERED AT 08:53:17 ON 11 MAR 2003)
                SET COST OFF
     FILE 'HCAPLUS' ENTERED AT 08:53:28 ON 11 MAR 2003
                E FUCOIDAN/CT
                E E3+ALL
L1
            684 S E4
     FILE 'REGISTRY' ENTERED AT 08:53:58 ON 11 MAR 2003
              1 S 9072-19-9
L2
L3
              5 S 9072-19-9/CRN
     FILE 'HCAPLUS' ENTERED AT 08:54:41 ON 11 MAR 2003
L4
           1028 S FUCOIDIN# OR FUCOIDAN# OR NEMACYSTUS MUCILAGE
L5
              5 S L3
           1048 S L1, L4, L5
L6
                E FUCOIDAN
            739 S E3
L.7
L8
            316 S E15, E17
L9
           1048 S L6-L8
L10
           225 S NITROGEN MONOOXIDE
     FILE 'REGISTRY' ENTERED AT 08:56:36 ON 11 MAR 2003
L11
              1 S 10102-43-9
     FILE 'HCAPLUS' ENTERED AT 08:57:40 ON 11 MAR 2003
          70393 S L11
T.12
L13
         135853 S OHM11771 OR OHM()(11771 OR 11 771) OR NITROGEN()(MONOXIDE OR
L14
              7 S L10, L12, L13 AND L9
                E INTERLEUKIN/CT
           5859 S E39
L15
                E E144+ALL
           6645 S E23,E46
L16
L17
          85930 S E7, E6+NT
         117789 S IL OR IL12 OR INTERLEUKIN OR (IL OR INTERLEUKIN) (L)12
L18
L19
             38 S L9 AND L15-L18
                E INTERFERON/CT
L20
            300 S E3
          30764 S E89
L21
                E E71+ALL
          54117 S E6+NT
L22
L23
          30764 S E6(L)GAMMA
          41617 S IFNGAMMA OR GAMMAIFN OR (IFN OR INTERFERON) (L) GAMMA
L24
             15 S L9 AND L20-L24
                E IGE/CT
                E E3+ALL
L26
           9782 S E2
                E IMMUNOGLOBULIN/CT
                E IMMUNOGLOBULINS/CT
```

```
10193 S E7, E6 (L) "E"
L28
              3 S L9 AND L26-L28
L29
               6 S L9 AND (IGE OR (IG OR IMMUNOGLOBULIN) (S) "E")
L30
                 E CYTOKINE/CT
                E E48+ALL
L31
          76929 S E5, E4
L32
         157526 S E4+NT
L33
              46 S L9 AND L31, L32
L34
              37 S L9 AND CYTOKINE
L35
              18 S L9 AND LYMPHOKINE
L36
              66 S L14, L19, L25, L29, L30, L33-L35
                E WO2000-JP5489/AP, PRN
               1 S E3, E4
L37
                E JP99-234262/AP, PRN
               1 S E4
L38
                E JP2000-69223/AP, PRN
               1 S E4
L39
                E TAKARA/PA,CS
            772 S E93-E129
L40
           1582 S E3-E145
L41
           3405 S (TAKARA? OR SHUZO?)/PA,CS
L42
              29 S L9 AND L40-L42
L43
                E TOMINAGA T/AU
           218 S E3, E4, E17-E19
L44
                 E TAKANARI/AU
                 E YAMASHITA S/AU
            385 S E3
L45
                E YAMASHITA SYU/AU
               7 S E6, E7
L46
                E SYUSAK/AU
                 E MIZUTANI S/AU
            106 S E3, E34
L47
                 E SHIGETOSHI/AU
                 E SAGAWA H/AU
L48
            386 S E3, E11, E12
                E HIROAKI S/AU
L49
               1 S E3
                E KATO I/AU
L50
            728 S E3-E5, E22-E25
                E IKUNOSH/AU
               5 S E4
L51
L52
              35 S L44-L51 AND L9
              2 S L36 AND L43, L52
L53
              34 S L43, L52 NOT L53
L54
              14 S L54 AND (FOOD# OR FEED? OR BEVERAGE# OR HEALTH FOOD# OR DRUG#
L55
                SEL DN AN 3 6 10 13 14
L56
               9 S L55 NOT E1-E15
              O S L54 AND (FOOD? OR NUTRI?)/SC, SX NOT L55
L57
             64 S L36 NOT L40-L57
L58
                 SEL DN AN 12 14 16 27 30 40 52
               7 S L58 AND E16-E36
L59
                 E ALLERGY/CT
                 E E3+ALL
L60
          19556 S E3, E2+NT
                 E E15+ALL
           7639 S E3
L61
                 E E7+ALL
L62
           6554 S E4
                 E E15+ALL
L63
          13841 S E5
                 E E4+ALL
          31196 S E4+NT
L64
                 E E13+ALL
```

```
L65
           8958 S E4, E3+NT
                E IMMUN/CT
                E IMMUNOS/CT
L66
          15342 S E12+NT OR E20+NT
          25407 S E26+NT OR E27+NT
L67
             26 S L9 AND L60-L67
L68
             11 S L68 NOT L36, L40-L59
L69
             91 S L9 AND (NUTRI? OR FOOD? OR FEED? OR BEVERAG? OR DRINK? OR JUI
L70
             79 S L9 AND (BEVERAG? OR ?DRINK? OR ?JUICE? OR FOOD? OR FEED?)/BI
L71
L72
             30 S L9 (L) FFD/RL
L73
             18 S L53, L56, L59
             4 S L68 AND L73
L74
L75
             18 S L73, L74
L76
             14 S L70-L72 AND L75
L77
             18 S L75, L76
L78
             89 S L70-L72 NOT L77
L79
             67 S L78 AND (PY<=2000 OR PRY<=2000 OR AY<=2000)
L80
             37 S L79 AND (FOOD? OR NUTRI?)/SC
                SEL DN AN 5 24
              2 S L80 AND E1-E6
L81
             20 S L77, L81
L82
L83
             30 S L79 NOT L80
             74 S L9 AND (?INFLAM? OR LEUKOTRIEN?)
L84
             19 S L84 AND L19, L29, L30, L36, L60-L69
L85
             1 S L84 AND L70-L72
L86
L87
             20 S L85, L86
                SEL DN AN 10 11 14 15 16 19 20
              7 S E7-E27 AND L87
L88
L89
             27 S L82, L88 AND L4-L10, L12-L88
```

FILE 'REGISTRY' ENTERED AT 09:53:35 ON 11 MAR 2003

=> fil hcaplus
FILE 'HCAPLUS' ENTERED AT 09:53:56 ON 11 MAR 2003
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 11 Mar 2003 VOL 138 ISS 11 FILE LAST UPDATED: 10 Mar 2003 (20030310/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

## => d 189 all hitstr tot

```
L89 ANSWER 1 OF 27 HCAPLUS COPYRIGHT 2003 ACS
AN 2002:800750 HCAPLUS
DN 137:293990
TI Functional foods, beverages, and feeds
containing fucoidan and agarooligosaccharides
```

```
ΙN
    Oyashiki, Haruo
PΑ
    Takara Bio Inc., Japan
    Jpn. Kokai Tokkyo Koho, 17 pp.
SO
    CODEN: JKXXAF
    Patent
DΤ
LA
    Japanese
IC
    ICM A23L001-308
         A21D002-18; A23C009-152; A23C011-10; A23C013-12; A23F003-14;
         A23G003-00; A23K001-16; A23L001-06; A23L001-16; A23L001-317;
          A23L001-325; A23L001-48; A23L002-52
CC
    17-6 (Food and Feed Chemistry)
    Section cross-reference(s): 18
FAN.CNT 1
                                                            DATE
    PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
     -----
                      ____
                                           -----
                            20021002
    JP 2002306131
                      A2
                                           JP 2001-117144
                                                            20010416
PΤ
PRAI JP 2001-117144
                            20010416
    Foods, beverages, or feeds contain
    fucoidan, its hydrolyzates, or their salts and
    agarooligosaccharides. The foods are effective for health
     improvement. Green tea beverage contg. fucoidan (from
    Kjellmaniella crassifolia) and agarooligosaccharides (prepd. from agar)
    was manufd.
ST
    food beverage feed fucoidan
    agarooligosaccharide; green tea fucoidan agarooligosaccharide
    health improvement
ΙT
    Pasta
        (Chinese; functional foods, beverages, and
        feeds contg. fucoidan and agarooligosaccharides)
IT
    Tea products
        (beverages, green; functional foods,
        beverages, and feeds contg. fucoidan and
        agarooligosaccharides)
IT
    Bakery products
        (buns; functional foods, beverages, and
        feeds contg. fucoidan and agarooligosaccharides)
TT
        (for fish or livestock; functional foods, beverages
        , and feeds contg. fucoidan and
        agarooligosaccharides)
TΤ
    Alcoholic beverages
    Bread
    Candy
    Chocolate
      Feed additives
      Food additives
    Milk
    Sake
    Soybean curd
        (functional foods, beverages, and feeds
        contg. fucoidan and agarooligosaccharides)
ΙT
    Oligosaccharides, biological studies
    RL: FFD (Food or feed use); SPN (Synthetic preparation); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (functional foods, beverages, and feeds
        contg. fucoidan and agarooligosaccharides)
IT
    Beverages
        (health; functional foods, beverages, and
        feeds contg. fucoidan and agarooligosaccharides)
IT
        (kamaboko; functional foods, beverages, and
        feeds contg. fucoidan and agarooligosaccharides)
ΙT
     Jams and Jellies
```

```
(orange; functional foods, beverages, and
        feeds contg. fucoidan and agarooligosaccharides)
TT
        (sausage; functional foods, beverages, and
        feeds contq. fucoidan and agarooligosaccharides)
IT
     Beverages
        (sports; functional foods, beverages, and
        feeds contg. fucoidan and agarooligosaccharides)
ΙT
     9002-18-0, Agar
     RL: CPS (Chemical process); PEP (Physical, engineering or chemical
     process); PROC (Process)
        (functional foods, beverages, and feeds
        contg. fucoidan and agarooligosaccharides)
IT
     9072-19-9P, Fucoidan
     RL: FFD (Food or feed use); PUR (Purification or recovery); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (functional foods, beverages, and feeds
        contg. fucoidan and agarooligosaccharides)
     5627-25-8P, Agarobiose
IT
     RL: FFD (Food or feed use); SPN (Synthetic preparation); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (functional foods, beverages, and feeds
        contg. fucoidan and agarooligosaccharides)
IT
     9072-19-9P, Fucoidan
     RL: FFD (Food or feed use); PUR (Purification or recovery); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (functional foods, beverages, and feeds
        contg. fucoidan and agarooligosaccharides)
     9072-19-9 HCAPLUS
RN
     Fucoidan (9CI)
                    (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     ANSWER 2 OF 27 HCAPLUS COPYRIGHT 2003 ACS
L89
     2002:220385 HCAPLUS
AN
DN
     136:252460
ΤI
     Homeostasis-maintaining agents
ΙN
     Nishiyama, Eiji; Sagawa, Hiroaki; Hino, Fumitsugu; Morihara,
     Etsuko; Sakai, Takeshi; Oyashiki, Haruo; Kato, Ikunoshin
PA
     Takara Shuzo Co., Ltd., Japan
SO
     PCT Int. Appl., 86 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     Japanese
     ICM A61K031-737
TC
         A61K035-80; A61P043-00; A61P001-16; A61P003-08; A61P003-06;
          A61P035-00; A61P031-18; A61P001-04; A23L002-52; A23L001-29;
          A23K001-16; C08B037-00
CC
     63-4 (Pharmaceuticals)
     Section cross-reference(s): 1, 17
                                                                             V.
FAN.CNT 1
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
                                                            DATE
                           _____
                                           ______
     ______
                      ____
                      A1
                            20020321
                                           WO 2001-JP7894
                                                            20/010912
PΙ
     WO 2002022140
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS,
             LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT,
             RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US,
             UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
```

```
20020326
    AU 2001088040
                       A5
                                           AU 2001-88040
                                                            20010912
PRAI JP 2000-278712
                       Α
                            20000913
    JP 2000-295077
                       Α
                            20000927
    JP 2000-342224
                      Α
                            20001109
    JP 2000-379313
                       Α
                            20001213
    JP 2001-128295
                      Α
                            20010425
    JP 2001-179335
                            20010613
                      Α
    WO 2001-JP7894
                       W
                            20010912
AB
    Disclosed are biol. homeostasis-maintaining agents, foods,
    drinks or feeds which comprise fucoidan, its
    decompn. products or salts thereof and have an effect of maintaining the
    homeostasis of living bodies. Also disclosed are fucoidan and
    marine algae exts. which are less colored, have relieved bitterness and a
     smaller iodine content and show a fresh feel; foods,
    drinks, seasonings, feeds, cosmetics or drugs contg. the
    above-mentioned fucoidan and/or marine algae exts.; and a
    process for efficiently producing the same.
ST
    fucoidan marine algae ext homeostasis food
ΙT
    Liver, disease
        (fibrosis; marine algae exts. as homeostasis-maintaining agents)
IT
    Cytoprotective agents
        (hepatoprotectants; marine algae exts. as homeostasis-maintaining
        agents)
IT
    Anti-AIDS agents
    Anticholesteremic agents
    Anticoagulants
    Antitumor agents
    Antiulcer agents
      Beverages
      Feed
      Food
    Homeostasis
    Kjellmaniella crassifolia
    Marine algae
    Seaweed
        (marine algae exts. as homeostasis-maintaining agents)
IT
    Lipids, biological studies
    RL: NPO (Natural product occurrence); PAC (Pharmacological activity); BIOL
     (Biological study); OCCU (Occurrence)
        (neutral, blood, lowering of; marine algae exts. as
        homeostasis-maintaining agents)
IΤ
     50-99-7, D-Glucose, biological studies
    RL: NPO (Natural product occurrence); PAC (Pharmacological activity); BIOL
     (Biological study); OCCU (Occurrence)
        (blood, lowering of; marine algae exts. as homeostasis-maintaining
        agents)
ΙT
     50-81-7, L-Ascorbic acid, biological studies
                                                    52-90-4, L-Cysteine,
    biological studies 70-18-8, Glutathione, biological studies
    Erythorbic acid
    RL: NPO (Natural product occurrence); PAC (Pharmacological activity); BIOL
     (Biological study); OCCU (Occurrence)
        (extn. in presence of reducing agents; marine algae exts. as
        homeostasis-maintaining agents)
                                         289890-05-7
IT
    9072-19-9, Fucoidan
                           184865-69-8
     289890-06-8
                   289890-07-9
                                 289890-08-0
                                              289890-09-1
                                                             289890-10-4
    RL: NPO (Natural product occurrence); PAC (Pharmacological activity); BIOL
     (Biological study); OCCU (Occurrence)
        (marine algae exts. as homeostasis-maintaining agents)
    9001-92-7, Protease
TT
    RL: NPO (Natural product occurrence); PAC (Pharmacological activity); BIOL
     (Biological study); OCCU (Occurrence)
        (pretreatment with; marine algae exts. as homeostasis-maintaining
        agents)
```

```
THERE ARE 46 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
        46
RE
(1) Kabushiki Kaisha Kibun Food Chemiphar; JP 6157519 A 1986
(2) Kabushiki Kaisha Kibun Food Chemiphar; JP 6157520 A 1986
(3) LI, D; Huazhong Nongye Daxue Xuebao 1999, V18(2), P191 HCAPLUS
(4) LI, F; Zhongguo Haiyang Yaowu 2000, V19(5), P12 HCAPLUS
(5) Takara Shuzo Co Ltd; EP 1057833 Al 1999 HCAPLÚS
(6) Takara Shuzo Co Ltd; JP 2001261704 A 1999 HCAPLUS
(7) Takara Shuzo Co Ltd; AU 9924401 Al 1999 HCAPLUS
(8) Takara Shuzo Co Ltd; WO 9941288 A1 1999 HCAPLUS
(9) Takara Shuzo Co Ltd; WO 0050464 Al 2000 HCAPLUS
(10) Takara Shuzo Co Ltd; WO 0062785 Al 2000 HCAPLUS
(11) Takara Shuzo Co Ltd; JP 2001224369 A 2000 HCAPLUS
(12) Takara Shuzo Co Ltd; JP 2001226392 A 2000 HCAPLUS
(13) Takara Shuzo Co Ltd; WO 0113925 A1 2001 HCAPLUS
(14) Takara Shuzoh Co Ltd; CN 1209749 A 1999 HCAPLUS
(15) Takara Shuzoh Co Ltd; CN 1221320 A 1999 HCAPLUS
(16) Takara Shuzoh Co Ltd; KR 2000010670 A 1999 HCAPLUS
(17) Takara Shuzoh Co Ltd; US 2001034335 A1 1999 HCAPLUS
(18) Takara Shuzoh Co Ltd; JP 2001218580 A 1999 HCAPLUS
(19) Takara Shuzoh Co Ltd; JP 2001224394 A2 1999 HCAPLUS
(20) Takara Shuzoh Co Ltd; JP 2001226407 A 1999 HCAPLUS
(21) Takara Shuzoh Co Ltd; JP 2001226408 A 1999 HCAPLUS
(22) Takara Shuzoh Co Ltd; CA 2243543 A 1999 HCAPLUS
(23) Takara Shuzoh Co Ltd; US 6207652 B1 1999 HCAPLUS
(24) Takara Shuzoh Co Ltd; AU 711896 B2 1999 HCAPLUS
(25) Takara Shuzoh Co Ltd; AU 720004 B2 1999 HCAPLUS
(26) Takara Shuzoh Co Ltd; EP 916269 Al 1999 HCAPLUS
(27) Takara Shuzoh Co Ltd; EP 919237 Al 1999 HCAPLUS
(28) Takara Shuzoh Co Ltd; AU 9673555 A1 1999
(29) Takara Shuzoh Co Ltd; AU 9713999 Al 1999 HCAPLUS
(30) Takara Shuzoh Co Ltd; WO 9726896 Al 1999 HCAPLUS
(31) Takara Shuzoh Co Ltd; AU 9727898 Al 1999 HCAPLUS
(32) Takara Shuzoh Co Ltd; WO 9747208 Al 1999 HCAPLUS
(33) Tanaka, M; JP 866159 A 1996
(34) Tanaka, Y; JP 01218573 A 1989
(35) Tanaka, Y; IT 1228458 A 1989
(36) Tanaka, Y; GB 2219484 A 1989
(37) Tanaka, Y; GB 2219484 B 1989
(38) Tanaka, Y; FR 2627672 A 1989
(39) Tanaka, Y; DE 3905866 A 1989
(40) Tanaka, Y; DE 3905866 C 1989
(41) Tanaka, Y; JP 475753 B 1989
(42) Tanaka, Y; US 4913915 A 1989
(43) Tanaka, Y; CH 681416 A 1989
(44) Tanaka, Y; IL 87261 A 1989
(45) Tanaka, Y; AU 8822122 A 1989
(46) Yu, K; Vopr Pitan 2000, V69(1/2), P22
TT
     9072-19-9, Fucoidan
     RL: NPO (Natural product occurrence); PAC (Pharmacological activity); BIOL
     (Biological study); OCCU (Occurrence)
        (marine algae exts. as homeostasis-maintaining agents)
     9072-19-9 HCAPLUS
RN
CN
     Fucoidan (9CI)
                     (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L89
     ANSWER 3 OF 27 HCAPLUS COPYRIGHT 2003 ACS
ΑN
     2001:254582 HCAPLUS
DN
     134:265601
     Foods containing fucoidan for treatment of NUD
TI
     (non-ulcer dyspepsia)
IN
     Yoshikawa, Masaki; Kudo, Tatsuyuki; Nagaoka, Masato; Hashimoto, Shusuke;
```

```
Kamiyama, Sadao; Shibata, Hideyuki; Takagi, Itsuko
     Yakult Honsha Co., Ltd., Japan
PΑ
     Jpn. Kokai Tokkyo Koho, 8 pp.
SO
     CODEN: JKXXAF
DT
     Patent
LA
     Japanese
IC
     ICM A23L001-30
     ICS A23L002-52; A23L002-38; A61K031-715; A61K035-78; A61K035-80;
         A61P001-04
CC
     17-13 (Food and Feed Chemistry)
FAN.CNT 1
                                          APPLICATION NO.
                     KIND DATE
                                                            DATE
     PATENT NO.
                      ____
                                           ------
     ______
                     A2
     JP 2001095528
                           20010410
                                           JP 1999-272232
                                                            19990927 <--
PΙ
PRAI JP 1999-272232
                           19990927 <--
     The foods contain fucoidan derived from brown algae.
     The foods, e.g., tea-like beverages, may also contain
     exts. from Senna tea, kaki leaves, Houttuynia cordata, and/or Foeniculum
     vulgare. A beverage contg. 100 mg fucoidan from
     Cladosiphon okamuranus improved gastric conditions in humans.
ST
     Cladosiphon fucoidan food nonulcer dyspepsia
     treatment; brown algae fucoidan beverage dyspepsia
     treatment
     Fennel (Foeniculum vulgare)
TΤ
     Houttuynia cordata
        (exts.; foods contg. brown algae fucoidan for
        treatment of NUD (nonulcer dyspepsia))
     Brown algae (Phaeophyceae)
IΤ
     Cladosiphon okamuranus
     Health food
        (foods contq. brown algae fucoidan for treatment of
        NUD (nonulcer dyspepsia))
TΥ
     Beverages
        (health; foods contg. brown algae fucoidan for
        treatment of NUD (nonulcer dyspepsia))
     Persimmon (Diospyros kaki)
T·T
        (leaf ext.; foods contg. brown algae fucoidan for
        treatment of NUD (nonulcer dyspepsia))
IT
     Dyspepsia
        (nonulcer; foods contg. brown algae fucoidan for
        treatment of NUD (nonulcer dyspepsia))
IT
     Senna (Cassia)
        (tea ext.; foods contg. brown algae fucoidan for
        treatment of NUD (nonulcer dyspepsia))
IT
     9072-19-9, Fucoidan
     RL: FFD (Food or feed use); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (foods contg. brown algae fucoidan for treatment of
        NUD (nonulcer dyspepsia))
TΤ
     9072-19-9, Fucoidan
     RL: FFD (Food or feed use); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (foods contg. brown algae fucoidan for treatment of
        NUD (nonulcer dyspepsia))
RN
     9072-19-9 HCAPLUS
CN
     Fucoidan (9CI)
                    (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L89
     ANSWER 4 OF 27 HCAPLUS COPYRIGHT 2003 ACS
AΝ
     2001:236061 HCAPLUS
DN
     135:151743
TΙ
     Functionality and health food application of seaweed
```

```
fucoidans
ΑU
     Sakai, Takeshi; Kato, Ikunoshi
     Bio Business Division, Takara Shuzo Co., Ltd., Japan
CS
     New Food Industry (2001), 43(2), 8-12
CODEN: NYFIAM; ISS 0547-0277
SO
PΒ
     Shokuhin Shizai Kenkyukai
     Journal; General Review
DT
LA
     Japanese
CC
     17-0 (Food and Feed Chemistry)
AΒ
     A review with 20 refs. on a no. of health foods and prodn. and
     usefulness thereof.
ST
     review health food seaweed fucoidan
ΤT
     Health food
     Seaweed
        (functionality and health food application of seaweed
        fucoidans)
     9072-19-9, fucoidan
TT
     RL: FFD (Food or feed use); BIOL (Biological study); USES (Uses)
        (functionality and health food application of seaweed
        fucoidans)
ΙT
     9072-19-9, fucoidan
     RL: FFD (Food or feed use); BIOL (Biological study); USES (Uses)
        (functionality and health food application of seaweed
        fucoidans)
     9072-19-9 HCAPLUS
RN
     Fucoidan (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    ANSWER 5 OF 27 HCAPLUS COPYRIGHT 2003 ACS
     2001:152496 HCAPLUS
AN
DN
     134:198038
TI
     Remedies containing fucoidan and/or its decomposition product
ΙN
     Tominaga, Takanari; Yamashita, Syusaku; Mizutani,
     Shigetoshi; Sagawa, Hiroaki; Kato, Ikunoshin
     Takara Shuzo Co., Ltd., Japan
PΑ
     PCT Int. Appl., 73 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     Japanese
TC.
     ICM A61K031-737
     ICS A61K035-80; A61K035-56; A61P037-02; A61P043-00; A61P037-08;
          C08B037-00
CC
     63-4 (Pharmaceuticals)
     Section cross-reference(s): 17
FAN.CNT 1
                                         APPLICATION NO. DATE
     PATENT NO.
                     KIND DATE
                     A1 20010301 WO 2000-JP5489 20000817 <--
     _____
     WO 2001013925
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
             HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU,
             LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD,
             SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU,
             ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
             CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                      AU 2000-65934
EP 2000-953450
                                                            20000817 <--
     AU 2000065934
                     A5 20010319
     EP 1226826
                      A1 20020731
                                                           20000817 <--
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL
PRAI JP 1999-234262 A 19990820 <--
```

```
JP 2000-69223
                            20000313
                       Α
                                      <--
                       W
                            20000817
                                      <--
     WO 2000-JP5489
     The invention relates to remedies or preventives for diseases with a need
AB
     for the regulation of the prodn. of cytokines, diseases with a
     need for the prodn. of nitrogen monoxide or allergic
     diseases characterized by contg. as the active ingredient {\it fucoidan}
     and/or its decompn. product; and foods, drinks or
     feeds for regulating the prodn. of cytokines,
     foods, drinks or feeds for inducing the prodn.
     of nitrogen monoxide, antiallergic foods,
     drinks or feeds, etc. contg. fucoidan and/or
     its decompn. product.
ST
     fucoidan cytokine regulation disease; antiallergy
     fucoidan decompn product; nitrogen monoxide
     disease fucoidan
TΤ
     Immunoglobulins
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (E, inhibitors; remedies contg. fucoidan and/or its
        decompn. product)
TI
     Algae
     Echinoderm (Echinodermata)
        (fucoidan from; remedies contg. fucoidan and/or its
        decompn. product)
ΙT
     Drug delivery systems
        (oral; remedies contg. fucoidan and/or its decompn. product)
IT
     Allergy inhibitors
       Beverages
       Feed
       Food
       Immunosuppressants
        (remedies contg. fucoidan and/or its decompn. product)
IT
     Cytokines
       Interferons
       Interleukin 12
       Interleukins
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (remedies contg. fucoidan and/or its decompn. product)
IT
     Interferons
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (.gamma.; remedies contg. fucoidan and/or its
        decompn. product)
TT
     10102-43-9, Nitrogen monoxide, biological
     studies
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (diseases related to prodn. of; remedies contg. fucoidan
        and/or its decompn. product)
     328081-45-4P
IT
     RL: PUR (Purification or recovery); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (remedies contg. fucoidan and/or its decompn. product)
IT
     9072-19-9, Fucoidan
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (remedies contg. fucoidan and/or its decompn. product)
RE.CNT
              THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD
        21
RE
(1) Dainippon Ink And Chemicals Inc; JP 09255577 A 1997 HCAPLUS
(2) Granert, C; Infect Immun 1999, V67(5), P2071 HCAPLUS
(3) Kyodo Nyugyo K K; JP 1072362 A 1998
(4) Shun, J; 1996, V14(3), P990 HCAPLUS
(5) The Australian National Universitay; JP 02502006 A
```

```
(6) The Australian National Universitay; JP 09328431 A HCAPLUS
(7) The Australian National Universitay; IL 106354 A1 HCAPLUS
(8) The Australian National Universitay; CA 1316828 A1 HCAPLUS
(9) The Australian National Universitay; AT 160941 E HCAPLUS
(10) The Australian National Universitay; AT 178212 E HCAPLUS
(11) The Australian National Universitay; JP 2701904 B2 HCAPLUS
(12) The Australian National Universitay; EP 355088 A1 HCAPLUS
(13) The Australian National Universitay; EP 355088 B1 HCAPLUS
(14) The Australian National Universitay; US 5541166 A HCAPLUS
(15) The Australian National Universitay; AU 605839 B2 HCAPLUS
(16) The Australian National Universitay; EP 631784 A1 HCAPLUS
(17) The Australian National Universitay; EP 631784 B1 HCAPLUS
(18) The Australian National Universitay; IL 85145 A1 HCAPLUS
(19) The Australian National Universitay; AU 8812410 A1 HCAPLUS
(20) The Australian National Universitay; WO 8805301 A1 1988 HCAPLUS
(21) Yokokawa, K; JOURNAL OF CLINICAL INVESTIGATION 1993, V92(4), P2080 HCAPLUS
IT
    10102-43-9, Nitrogen monoxide, biological
    studies
    RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
    BIOL (Biological study); OCCU (Occurrence)
        (diseases related to prodn. of; remedies contg. fucoidan
        and/or its decompn. product)
RN
     10102-43-9 HCAPLUS
CN
    Nitrogen oxide (NO) (8CI, 9CI) (CA INDEX NAME)
N = 0
     9072-19-9, Fucoidan
IT
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (remedies contq. fucoidan and/or its decompn. product)
RN
     9072-19-9 HCAPLUS
     Fucoidan (9CI)
                     (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
T.89
    ANSWER 6 OF 27 HCAPLUS COPYRIGHT 2003 ACS
ΑN
    2000:756531 HCAPLUS
DN
    133:325613
    Remedies or preventives for diseases with need for growth factor
TΙ
    production-inducing effect
IN
    Sagawa, Hiroaki; Sakai, Takeshi; Kobayashi, Eiji; Li, Tuo-Ping;
    Ohnogi, Hiromu; Nishimura, Kaori; Nishiyama, Eiji; Wu, Hua-Kang;
    Mizutani, Shigetoshi; Kato, Ikunoshin
PA
    Takara Shuzo Co., Ltd., Japan
    PCT Int. Appl., 158 pp.
SO
    CODEN: PIXXD2
DТ
    Patent
LA
    Japanese
    A61K031-737; A61K031-70; A61K031-7016; A61K031-702; A61K007-40;
TC
    A61P043-00; A23L002-52; A23L001-29; A23K001-16; C08B037-00
CC
    63-4 (Pharmaceuticals)
    Section cross-reference(s): 17, 62
FAN.CNT 2
                      KIND DATE
                                           APPLICATION NO.
    PATENT NO.
                                                            DATE
                                           -----
     ------
                           (20001026)
PΙ
    WO 2000062785
                      A1
                                           WO 2000-JP2432
                                                            20000414
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR,
             CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
             ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV,
             MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG,
             SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW,
```

```
AM, AZ, BY, KG, KZ, MD, RU; TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
             DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
             CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     EP 1175907
                       Α1
                            20020130
                                           EP 2000-917309
                                                             20000414
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO
                            19990415
PRAI JP 1999-108067
                       Α
     JP 1999-108499
                            19990415
                       Α
     JP 1999-114542
                       Α
                            19990422
     JP 1999-129163
                            19990510
                       Α
     JP 1999-142343
                       Α
                            19990521
     JP 1999-154662
                       Α
                            19990602
     JP 1999-200982
                       Α
                            19990714
     JP 1999-275231
                       Α
                            19990928
     JP 1999-375606
                       Α
                            19991228
     JP 2000-99941
                       Α
                            20000331
    WO 2000-JP2432
                       W
                            20000414
    MARPAT 133:325613
OS
     The invention relates to remedies or preventives for diseases with a need
AB
     for a growth factor prodn.-inducing effect, characterized by contg.
    member(s) selected from the group consisting of acidic polysaccharides and
     degrdn. products thereof, acidic oligosaccharides, acidic monosaccharides,
     acidic sugar alcs. and salts thereof each having an effect of inducing the
     prodn. of growth factor; foods, drinks or
     feeds for inducing the prodn. of growth factor; cosmetics for
     inducing the prodn. of growth factor; and growth factor prodn. regulators.
ST
     acidic polysaccharide disease growth factor induction
    Alditols
ТТ
    Monosaccharides
    Oligosaccharides, biological studies
     Polysaccharides, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (acidic; remedies or preventives for diseases with need for growth
        factor prodn.-inducing effect)
TΤ
    Cosmetics
        (cleansing, facial; remedies or preventives for diseases with need for
        growth factor prodn.-inducing effect)
TΤ
    Cosmetics
        (creams; remedies or preventives for diseases with need for growth
        factor prodn.-inducing effect)
IT
    Cosmetics
        (emulsions; remedies or preventives for diseases with need for growth
        factor prodn.-inducing effect)
IT
     Cosmetics
        (lotions; remedies or preventives for diseases with need for growth
        factor prodn.-inducing effect)
ΙT
    Cosmetics
        (packs; remedies or preventives for diseases with need for growth
        factor prodn.-inducing effect)
ΙT
    Algae
    Animal
    Bath preparations
       Beverages
     Cosmetics
     Detergents
     Disease, animal
       Feed
     Fish
       Food
    Microorganism
     Plant (Embryophyta)
```

(remedies or preventives for diseases with need for growth factor

prodn.-inducing effect)

IT Growth factors, animal

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence) (remedies or preventives for diseases with need for growth factor prodn.-inducing effect)

IT Soaps

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(remedies or preventives for diseases with need for growth factor prodn.-inducing effect)

IT Cytokines

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (remedies or preventives for diseases with need for growth factor prodn.-inducing effect)

IT Prostaglandins

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (remedies or preventives for diseases with need for growth factor prodn.-inducing effect)

IT Polysaccharides, biological studies

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (sulfated; remedies or preventives for diseases with need for growth factor prodn.-inducing effect)

IT 9061-61-4, Nerve growth factor 49557-75-7, Liver cell growth factor 61912-98-9, Insulin-like growth factor

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence)

(remedies or preventives for diseases with need for growth factor

(remedies or preventives for diseases with need for growth factor prodn.-inducing effect)

58-86-6D, Xylose, sulfated 59-23-4D, ΙT 57-50-1D, Sucrose, sulfated Galactose, sulfated 63-42-3D, Lactose, sulfated 69-79-4D, Maltose, 99-20-7D, Trehalose, sulfated 154-17-6D, 2-DeoxyGlucose, sulfated 287-92-3, Cyclopentane 499-40-1D, IsoMaltose, sulfated 547-25-1D, Turanose, sulfated 528-50-7D, Cellobiose, sulfated 1109-28-0D, Maltotriose, sulfated 585-99-9D, Melibiose, sulfated 3458-28-4D, Mannose, sulfated 4618-18-2D, Lactulose, sulfated **9072-19-9**, **Fucoidan** 13718-94-0D, Palatinose, sulfated 30077-17-9D, Talose, sulfated 33038-63-0, Glucose sulfate 34620-77-4D, 34620-77-4D, Maltohexaose, sulfated Maltohexaose, dodecyl, sulfated 34620-78-5D, Maltoheptaose, sulfated

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (remedies or preventives for diseases with need for growth factor prodn.-inducing effect)

RE.CNT 30 THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

- (1) Amrad Operations Pty Ltd; WO 9623003 A1 1996 HCAPLUS
- (2) Belford; J Cell Physiol 1993, V157(1), P184 HCAPLUS
- (3) Cancer Research Campaign Technology Limited; JP 07507596 A
- (4) Cancer Research Campaign Technology Limited; EP 642533 A1 HCAPLUS
- (5) Cancer Research Campaign Technology Limited; WO 9421689 Al 1994 HCAPLUS
- (6) Glycomed Incorporated; JP 09503510 A
- (7) Glycomed Incorporated; US 5739115 A HCAPLUS
- (8) Glycomed Incorporated; EP 722326 Al HCAPLUS
- (9) Glycomed Incorporated; WO 9509637 A1 1995 HCAPLUS
- (10) Glycomed Incorporated; WO 9609828 Al 1996 HCAPLUS
- (11) Mendes; WO 9712598 A1 1997 HCAPLUS
- (12) Mendes; WO 9712598 A1 1997 HCAPLUS
- (13) Mitsubishi Kasei Corp; JP 05301824 A 1993 HCAPLUS
- (14) Mitsui Norin K K; JP 07173059 A 1995 HCAPLUS
- (15) Nakamura; US 5977310 A HCAPLUS
- (16) Nakamura; WO 9628475 A1 HCAPLUS

```
(17) Nakamura; EP 816381 Al 1998 HCAPLUS
(18) Nakamura, T; JP 06312941 A 1994 HCAPLUS
(19) Prestrelski; Arch Biochem Biophys 1992, V293(2), P314 HCAPLUS
(20) Snow Brand Milk Products Co Ltd; JP 07267995 A HCAPLUS
(21) Snow Brand Milk Products Co Ltd; WO 9526984 A1 1995 HCAPLUS
(22) Takara Shuzo Co Ltd; EP 919237 A1 HCAPLUS
(23) Takara Shuzo Co Ltd; EP 941981 A1 HCAPLUS
(24) Takara Shuzo Co Ltd; EP 976717 Al HCAPLUS
(25) Takara Shuzo Co Ltd; EP 984001 Al HCAPLUS
(26) Takara Shuzo Co Ltd; WO 9726896 A1 1997 HCAPLUS
(27) Takara Shuzo Co Ltd; WO 9813328 Al 1998 HCAPLUS
(28) Takara Shuzo Co Ltd; WO 9839291 A1 1998 HCAPLUS
(29) Takara Shuzo Co Ltd; WO 9840346 Al 1998 HCAPLUS
(30) Takara Shuzo Co Ltd; WO 9941288 A1 1999 HCAPLUS
TΨ
     9072-19-9, Fucoidan
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (remedies or preventives for diseases with need for growth factor
        prodn.-inducing effect)
     9072-19-9 HCAPLUS
RN
     Fucoidan (9CI)
                     (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     ANSWER 7 OF 27 HCAPLUS COPYRIGHT 2003 ACS
L89
ΑN
     2000:754801 HCAPLUS
DN
     134:192344
ΤI
     Functionality and effects of fucoidans
     Kato, Ikunoshin; Sakai, Takeshi; Sagawa, Hiroaki
ΑU
CS
     Bio Reaearch Lab., Takara Shuzo Co., Japan
SO
     Japan Fudo Saiensu (2000), 39(9), 43-47
     CODEN: JAFSAA; ISSN: 0368-1122
PB
     Nippon Shokuhin Shuppan K.K.
\mathsf{DT}
     Journal; General Review
LA
     Japanese
CC
     17-0 (Food and Feed Chemistry)
AB
     A review with 10 refs. on the chem. structures of fucoidans and
     usefulness of these compds. for health foods.
ST
     review fucoidan chem structure health food
TT
     Health food
        (functionality and effects of fucoidans for)
ΙT
     Molecular structure
        (functionality and effects of fucoidans in relation to)
ΙT
     9072-19-9, Fucoidan 9072-19-9D,
     Fucoidan, analogs
     RL: FFD (Food or feed use); BIOL (Biological study); USES (Uses)
        (functionality and effects of fucoidans)
IT
     9072-19-9, Fucoidan 9072-19-9D,
     Fucoidan, analogs
     RL: FFD (Food or feed use); BIOL (Biological study); USES (Uses)
        (functionality and effects of fucoidans)
     9072-19-9 HCAPLUS
RN
CN
     Fucoidan (9CI)
                    (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
RN
     9072-19-9 HCAPLUS
     Fucoidan (9CI)
                    (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L89
     ANSWER 8 OF 27 HCAPLUS COPYRIGHT 2003 ACS
     2000:358976 HCAPLUS
AN
DN
     133:103638
     Inhibition of leukocyte entry into the brain by the selectin blocker
TΙ
```

fonda 0 10 / 049419 fucoidin decreases interleukin-1 (IL-1) levels but increases IL-8 levels in cerebrospinal fluid during experimental pneumococcal meningitis in rabbits Ostergaard, Christian; Yieng-Kow, Runa Vavia; Benfield, Thomas; Frimodt-Moller, Niels; Espersen, Frank; Lundgren, Jens D. Division of Microbiology, Department of Research and Development, Statens Serum Institut, Copenhagen DK-2300, Den. Infection and Immunity (2000), 68(6), 3153-3157 CODEN: INFIBR; ISSN: 0019-9-67 American Society for Microbiology Journal English 15-8 (Immunochemistry) The polysaccharide fucoidin is a selectin blocker that inhibits leukocyte recruitment into the cerebrospinal fluid (CSF) during exptl. pneumococcal meningitis. In the present study, the effect of fucoidin treatment on the release of the proinflammatory cytokines tumor necrosis factor alpha (TNF-.alpha.), interleukin-1 (IL-1), and IL-8 into the CSF was investigated. Rabbits (n = 7) were treated i.v. with 10 mg of fucoidin/kg of body wt. every second hour starting 4 h after intracisternal inoculation of .apprx.106 CFU of Streptococcus pneumoniae type 3 (untreated control group, n = 7). CSF samples were obtained every second hour during a 16-h study period. Treatment with fucoidin caused a consistent and significant decrease in CSF IL-1 levels (in picograms per mL) between 12 and 16 h (0 vs. 170, 0 vs. 526, and 60 vs. 1,467, resp.; P < 0.02). A less consistent decrease in CSF TNF-.alpha. levels was obsd. in the fucoidin-treated group, but with no significant difference between the two groups (P > 0.05). contrast, there was no attenuation in CSF IL-8 levels. Indeed, there was a significant increase in CSF IL-8 levels (in picograms per mL) in the fucoidin-treated group at 10 and **12** h (921 vs. 574 and 1,397 vs. 569, resp.; P < 0.09). In conclusion, our results suggest that blood-derived leukocytes mainly are responsible for the release of IL-1 and to some degree TNF-.alpha. into the CSF during pneumococcal meningitis, whereas IL-8 may be produced by local cells within the brain. leukocyte selectin fucoidin interleukin cerebrospinal fluid pneumococcal meningitis Meningitis (bacterial; inhibition of leukocyte entry into brain by selectin blocker fucoidin decreases interleukin-1 but increases IL-8 in cerebrospinal fluid during pneumococcal meningitis in rabbits) Brain

IT

ST

IT

ΑU

CS

SO

PB

 $\mathsf{DT}$ 

LA

CC

AB

Cerebrospinal fluid

Streptococcus pneumoniae

(inhibition of leukocyte entry into brain by selectin blocker fucoidin decreases interleukin-1 but increases IL-8 in cerebrospinal fluid during pneumococcal meningitis in rabbits)

ΙT Selectins

> RL: ADV (Adverse effect, including toxicity); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(inhibition of leukocyte entry into brain by selectin blocker fucoidin decreases interleukin-1 but increases IL-8 in cerebrospinal fluid during pneumococcal meningitis in rabbits)

ΙT Interleukin 1

RL: BPR (Biological process); BSU (Biological study, unclassified); MFM (Metabolic formation); BIOL (Biological study); FORM (Formation, nonpreparative); PROC (Process)

(inhibition of leukocyte entry into brain by selectin blocker fucoidin decreases interleukin-1 but increases IL-8 in cerebrospinal fluid during pneumococcal meningitis in rabbits)

## IT Interleukin 8

RL: BPR (Biological process); BSU (Biological study, unclassified); MFM (Metabolic formation); BIOL (Biological study); FORM (Formation, nonpreparative); PROC (Process)

(inhibition of leukocyte entry into brain by selectin blocker **fucoidin** decreases **interleukin-1** but increases **IL-8** in cerebrospinal fluid during pneumococcal meningitis in rabbits)

IT Tumor necrosis factors

RL: BPR (Biological process); BSU (Biological study, unclassified); MFM (Metabolic formation); BIOL (Biological study); FORM (Formation, nonpreparative); PROC (Process)

(inhibition of leukocyte entry into brain by selectin blocker fucoidin decreases interleukin-1 but increases IL-8 in cerebrospinal fluid during pneumococcal meningitis in

IL-8 in cerebrospinal fluid during pneumococcal meningitis rabbits)

IT Blood-brain barrier

(inhibition of leukocyte entry into brain by selectin blocker fucoidin decreases interleukin-1 but increases

IL-8 in cerebrospinal fluid during pneumococcal meningitis in rabbits in relation to)

IT Cell migration

(leukocyte infiltration; inhibition of leukocyte entry into brain by selectin blocker fucoidin decreases interleukin-1 but increases IL-8 in cerebrospinal fluid during pneumococcal meningitis in rabbits)

IT 9072-19-9, Fucoidin

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(inhibition of leukocyte entry into brain by selectin blocker fucoidin decreases interleukin-1 but increases IL-8 in cerebrospinal fluid during pneumococcal meningitis in rabbits)

RE.CNT 26 THERE ARE 26 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

- (1) Angstwurm, K; Neurosci Lett 1995, V191, P1 HCAPLUS
- (2) Bitsch, A; Neurosci Lett 1997, V237, P105 HCAPLUS
- (3) Burroughs, M; Microb Pathog 1995, V19, P245 HCAPLUS
- (4) Dacey, R; Antimicrob Agents Chemother 1974, V6, P437 HCAPLUS
- (5) Engelhard, D; J Infect Dis 1997, V175, P979 MEDLINE
- (6) Friedland, I; J Infect Dis 1995, V172, P805 HCAPLUS
- (7) Granert, C; Infect Immun 1999, V67, P2071 HCAPLUS
- (8) Granert, C; J Clin Investig 1994, V93, P929 HCAPLUS
- (9) Harada, A; J Leukoc Biol 1994, V56, P559 HCAPLUS
- (10) Lowry, O; J Biol Chem 1951, V193, P265 HCAPLUS
- (11) Matsukawa, A; Inflamm Res 1998, V47, PS137 HCAPLUS
- (12) Molvig, J; Scand J Immunol 1987, V26, P611 HCAPLUS
- (13) Mustafa, M; J Clin Investig 1989, V84, P1253 HCAPLUS
- (14) Nau, R; J Antimicrob Chemother 1997, V39, P781 HCAPLUS
- (15) Ostergaard, C; Antimicrob Agents Chemother 1998, V42, P1706 HCAPLUS
- (16) Ostergaard, C; Eur J Clin Microbiol Infect Dis 1996, V15, P166 MEDLINE
- (17) Ostergaard, C; Infect Immun 1999, V67, P3430 HCAPLUS
- (18) Quagliarello, V; J Clin Investig 1991, V87, P1360 HCAPLUS
- (19) Ramilo, O; J Exp Med 1990, V172, P497 HCAPLUS
- (20) Saez-Llorens, X; J Clin Investig 1991, V88, P2003 HCAPLUS
- (21) Saukkonen, K; J Exp Med 1990, V171, P439 HCAPLUS
- (22) Spector, R; J Clin Investig 1974, V54, P316 HCAPLUS
- (23) Tang, T; J Clin Investig 1996, V97, P2485 HCAPLUS

- (24) Tauber, M; Clin Infect Dis 1999, V28, P1 HCAPLUS
- (25) Waage, A; J Exp Med 1989, V170, P1859 HCAPLUS
- (26) Zysk, G; J Neuroimmunol 1997, V73, P77 HCAPLUS
- IT 9072-19-9, Fucoidin

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(inhibition of leukocyte entry into brain by selectin blocker **fucoidin** decreases **interleukin**-1 but increases **IL-8** in cerebrospinal fluid during pneumococcal meningitis in rabbits)

- RN 9072-19-9 HCAPLUS
- CN Fucoidan (9CI) (CA INDEX NAME)
- \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*
- L89 ANSWER 9 OF 27 HCAPLUS COPYRIGHT 2003 ACS
- AN 2000:250708 HCAPLUS
- DN 133:115050
- TI Effects of sea tangle (Laminaria japonica) extract and **fucoidan drinks** on oxygen radicals and their scavenger enzymes in stressed
  mouse
- AU Choi, Jin-Ho; Kim, Dae-Ik; Park, Soo-Hyun; Kim, Dong-Woo; Kim, Chang Mok; Koo, Jae Geun
- CS Lab. of Biochemistry, Pukyong National University, Pusan, 608-737, S. Korea
- SO Han'guk Susan Hakhoechi (1999), 32(6), 764-769 CODEN: HSHKAW; ISSN: 0374-8111
- PB Korean Fisheries Society
- DT Journal
- LA English
- CC 1-12 (Pharmacology)
- This study was designed to investigate the effects of sea tangle AB (Laminaria japonica) ext. (Dasi-Ex group: dry base 4.0%) and fucoidan-added (Fuco-I, II, III group: fucoidan of 1.0%, 2.0%, 3.0% added to Dasi-Ex) drinks on the formation of oxygen radicals and scavenger enzyme activities of stressed mice. ICR male mice (20 g) were fed exptl. diets and these drinks instead of water for 18 days including 4 days of sociopsychol. stress. Dasi-Ex and Fuco-I, II and III groups resulted in a marked decreases 20.apprx.40% in basal oxygen radical (BOR) formation, and 15.apprx.25% in induced oxygen radical (IOR) formation compared with control group. Hydroxyl radical formations were significantly inhibited about 10% in Dasi-Ex group, while remarkably inhibited 30.apprx.40% in Fuco-I, II and III groups. Lipid peroxide (LPO) levels in Dasi-Ex group were not significantly different from those of control group, but Fuco-I, II and III groups resulted in a significant decreases about 10% in LPO levels compared with control group. Dasi-Ex, Fuco-I, II and III groups resulted in a marked decreases (31%, 36%, 39% and 42%, resp.) in oxidized protein levels through prodn. of carbonyl group. Significant differences in nitric oxide (NO) levels in Dasi-Ex group were not obtained, but NO levels were slightly inhibited about 7% in Fuco-I and II groups and 20% in Fuco-III group compared with control group. Significant differences in superoxide dismutase (SOD) and catalase (CAT) activities in Dasi-Ex and Fuco-I groups were not obtained, but Fuco-II and III groups resulted in a significant increases 25.apprx.40% in SOD activities, and about 10% in CAT activities compared with control group. These results suggest that the sociopsychol. stress and aging process could be effectively inhibited by biol. activity of sea tangle and fucoidan components.
- ST sea tangle fucoidan oxygen radical stress
- IT Laminaria japonica

(effects of sea tangle (Laminaria japonica) ext. and **fucoidan** drinks on oxygen radicals and scavenger enzymes in stressed

7 date

mouse) Reactive oxygen species ΙT RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (effects of sea tangle (Laminaria japonica) ext. and fucoidan drinks on oxygen radicals and scavenger enzymes in stressed mouse) Peroxides, biological studies IT Peroxides, biological studies RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (lipid; effects of sea tangle (Laminaria japonica) ext. and fucoidan drinks on oxygen radicals and scavenger enzymes in stressed mouse) TT Proteins, general, biological studies RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (oxidized; effects of sea tangle (Laminaria japonica) ext. and fucoidan drinks on oxygen radicals and scavenger enzymes in stressed mouse) IT Lipids, biological studies Lipids, biological studies RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (peroxides; effects of sea tangle (Laminaria japonica) ext. and fucoidan drinks on oxygen radicals and scavenger enzymes in stressed mouse) IΤ Antioxidants (pharmaceutical; effects of sea tangle (Laminaria japonica) ext. and fucoidan drinks on oxygen radicals and scavenger enzymes in stressed mouse) ΙT Stress, animal (psychosocial; effects of sea tangle (Laminaria japonica) ext. and fucoidan drinks on oxygen radicals and scavenger enzymes in stressed mouse) TT 9072-19-9, Fucoidan RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (effects of sea tangle (Laminaria japonica) ext. and fucoidan drinks on oxygen radicals and scavenger enzymes in stressed mouse) ΙΤ 3352-57-6, Hydroxyl radical, biological studies 7782-44-7D, Oxygen, 9001-05-2, Catalase 9054-89-1, reactive species, biological studies Superoxide dismutase 10102-43-9, Nitric oxide , biological studies RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (effects of sea tangle (Laminaria japonica) ext. and fucoidan drinks on oxygen radicals and scavenger enzymes in stressed mouse) RE.CNT 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD RE (1) Applegate, E; J Nutr 1984, V114, P447 HCAPLUS (2) Bernardi, G; J Biol Chem 1962, V273, P75 (3) Choi, J; Age 1989, V12, P133 HCAPLUS (4) Choi, J; Age 1990, V13, P61 HCAPLUS (5) Choi, J; Age 1996, V19, P1 HCAPLUS (6) Choi, J; Age & Nutrition 1999, V10(1), P47 (7) Choi, J; Free Rad Biol & Med 1995, V18(2), P133 HCAPLUS (8) Choi, J; J Biochem 1991, V23(1), P61 HCAPLUS (9) Choi, J; J Kor Soc Food Nutr 1991, V20(2), P187 HCAPLUS (10) Choi, J; J Nutr Health & Aging 1998, V2(3), P451

```
(11) Choi, J; J Nutr Health & Aging 1998, V2(3), P456
 (12) Choi, J; J Nutr Health & Aging 1998, V2(3), P461
 (13) Choi, J; Kor J Gerontol 1994, V4(2), P61
 (14) Choi, J; Kor J Life Sci 1998, V9(5), P604
(15) Choi, J; Kor J Life Sci 1998, V9(5), P612

(16) Choi, J; Kor J Life Sci 1998, V9(5), P612

(17) Choi, J; Kor J Life Sci 1999, V9(4), P430

(18) Choi, J; Kor J Life Sci 1999, V9(4), P430

(19) Choi, J; Kor J Life Sci 1999, V9(4), P430

(20) Choi, J; Kor J Life Sci 1999, V9(5), P537

(21) The H: Chom Pharm Pull 1976, V2(4), P114
 (21) Ito, H; Chem Pharm Bull 1976, V24, P114
 (22) Lowry, O; J Biol Chem 1951, V193, P265 HCAPLUS
 (23) Nakazawa, Y; Chemotherapy 1974, V22, P1435
 (24) Ogawa, N; Jpn J Psychosom Med 1966, V6, P352
 (25) Oyanagui, Y; Anal Biochem 1984, V42, P290
 (26) Rigo, A; Anal Biochem 1977, V81, P157 HCAPLUS
 (27) Usui, T; Agric Biol Chem 1980, V44, P1965 HCAPLUS
 (28) Yagi, K; Chemistry and Physics of Lipids 1987, V45, P337 HCAPLUS
 (29) Yamamoto, I; J Exp Med 1981, V51, P187 MEDLINE (30) Yu, B; Ann NY Acad Sci 1996, V786, P1 HCAPLUS
.(31) Yu, B; Free Rad Biol Med 1996, V21, P651 HCAPLUS
 (32) Yu, B; Soc Exp Biol Med 1990, V193, P13 HCAPLUS
      9072-19-9, Fucoidan
      RL: BAC (Biological activity or effector, except adverse); BSU (Biological
      study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
       (Uses)
          (effects of sea tangle (Laminaria japonica) ext. and fucoidan
          drinks on oxygen radicals and scavenger enzymes in stressed
          mouse)
      9072-19-9
 RN
                   HCAPLUS
      Fucoidan (9CI) (CA INDEX NAME)
CN
 *** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
      10102-43-9, Nitric oxide, biological studies
 TT
      RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
       (Biological study); PROC (Process)
          (effects of sea tangle (Laminaria japonica) ext. and fucoidan
          drinks on oxygen radicals and scavenger enzymes in stressed
          mouse)
      10102-43-9 HCAPLUS
RN
      Nitrogen oxide (NO) (8CI, 9CI) (CA INDEX NAME)
. CN
N = 0
      ANSWER 10 OF 27 HCAPLUS COPYRIGHT 2003 ACS
 1.89
 AN
      2000:111177 HCAPLUS
      132:121786
 DN
 ΤI
      Food and pharmaceuticals containing microbicidal
      fucoidan
      Sakai, Takeshi; Kimura, Hitomi; Katayama, Kaoru; Kato, Ikunoshin
 ΙN
 PA
      Takara Shuzo Co., Ltd., Japan
 SO
      Jpn. Kokai Tokkyo Koho, 5 pp.
      CODEN: JKXXAF
 DT
      Patent
 LA
      Japanese
 IC
      ICM C08B037-00
      ICS A23L001-30; A23L002-52; A23L002-38; A61K031-725; A61K035-80
 CC
      17-6 (Food and Feed Chemistry)
      Section cross-reference(s): 63
```

FAN.CNT 1

```
PATENT NO.
                     KIND DATE
                                          APPLICATION NO.
                                                           DATE
                     ----
                                          _____
    ______
                                                           -----
                    A2 20000215
    JP 2000044602
                                          JP 1998-217282
                                                           19980731
                                                                          7,000
                     . 19980731
PRAI JP 1998-217282
    Fucoidan and/or derivs. as microbicides against Helicobacter,
    are added to food, beverages, and pharmaceuticals.
    microbicide Helicobacter food beverage pharmaceutical
ST
IT
    Helicobacter
        (food and pharmaceuticals contg. microbicidal
       fucoidan against)
ΙT
    Beverages
      Drugs
       Food
        (microbicidal fucoidan for)
IT
    9072-19-9, Fucoidan 9072-19-9D,
    Fucoidan, derivs.
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); FFD (Food or feed use); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (food and pharmaceuticals contg. microbicidal
        fucoidan)
IT
    9072-19-9, Fucoidan 9072-19-9D,
    Fucoidan, derivs.
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
    study, unclassified); FFD (Food or feed use); THU (Therapeutic
    use); BIOL (Biological study); USES (Uses)
        (food and pharmaceuticals contg. microbicidal
        fucoidan)
RN
     9072-19-9 HCAPLUS
CN
    Fucoidan (9CI) (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     9072-19-9 HCAPLUS
RN
CN
    Fucoidan (9CI) (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L89 ANSWER 11 OF 27 HCAPLUS COPYRIGHT 2003 ACS
AN
    1999:816002 HCAPLUS
DN
    132:63283
TΙ
    Characteristics and utilization of "fucoidans" from seaweeds and
    their oligosaccharides
                                                                       7 date
ΑU
    Sakai, Takeshi; Kato, Ikunoshin
    Bio-Hirosaki Res. Lab., Takara Shuzo Co., Ltd., Japan
CS .
SO
    Gekkan Fudo Kemikaru (1999), 15(12), 66-71
    CODEN: GFKEEX; ISSN: 0911-2286
PΒ
    Shokuhin Kagaku Shinbunsha
DT
    Journal; General Review
LA
    Japanese
CC
    17-0 (Food and Feed Chemistry)
    Section cross-reference(s): 62
    A review with 26 refs., on characterization and application of
AB
    fucoidan, which is a group of polysaccharides contg. sulfated
     fucose, from seaweed, discussing prepn. methods, discovery of
    fucoidan-degrading enzymes and their application, physicochem.
    properties, function, safety, and food or cosmetic application.
ST
    review fucoidan oligosaccharide seaweed food cosmetic
IT
    Cosmetics
       Food
     Seaweed
        (characteristics and utilization of fucoidans from seaweeds
        and their oligosaccharides)
```

ΙT

Oligosaccharides, properties

```
RL: PRP (Properties)
        (characteristics and utilization of fucoidans from seaweeds
        and their oligosaccharides)
     Polysaccharides, biological studies
IT
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BUU (Biological use, unclassified); FFD (Food
     or feed use); PRP (Properties); PUR (Purification or recovery); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (fucoidans; characteristics and utilization of
        fucoidans from seaweeds and their oligosaccharides)
ΙT
     9072-19-9P, Fucoidan
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BUU (Biological use, unclassified); FFD (Food
     or feed use); PRP (Properties); PUR (Purification or recovery); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (characteristics and utilization of fucoidans from seaweeds
        and their oligosaccharides)
ΙT
     9072-19-9P, Fucoidan
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BUU (Biological use, unclassified); FFD (Food
     or feed use); PRP (Properties); PUR (Purification or recovery); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (characteristics and utilization of fucoidans from seaweeds
        and their oligosaccharides)
     9072-19-9 HCAPLUS
     Fucoidan (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L89 ANSWER 12 OF 27 HCAPLUS COPYRIGHT 2003 ACS
AN
     1999:531032 HCAPLUS
DN
     131:183861
TI
     Immunity-enhancing agent and food
ΙN
     Hori, Tetsuji; Kiyoshima, Junko; Yasui, Hisako
     Yakult Honsha Co., Ltd., Japan
PΑ
     Jpn. Kokai Tokkyo Koho, 4 pp.
SO
                                                                        7 date
     CODEN: JKXXAF
DT
     Patent
LA
     Japanese
IC
     ICM C08B037-00
     ICS A23L001-30; A61K031-725; A61K035-80
     15-2 (Immunochemistry)
     Section cross-reference(s): 1, 6, 17
FAN.CNT 1
     PATENT NO.
                    KIND DATE
                                          APPLICATION NO. DATE
     ______
                                          -----
PI JP 11228602 A2 19990824.
PRAI JP 1998-41043 19980209
                                          JP 1998-41043
                                                          19980209
     Fucoidan, a sulfated polysaccharide, is disclosed as an enhancer
     for humoral and cellular immunity. Also disclosed are safe, low cost and
     good taste food products contg. fucoidan as effective
     ingredient in enhancing immunity. Fucoidan was purified from
     seaweed of Spermatochnaceae, and used for inducing prodn. of antibody
     (i.e. IgA, IgM and IgG) and interferon-.gamma...
ST
     humoral cellular immunity enhancer fucoidan food
ΙT
     Immunoglobulins
     RL: BSU (Biological study, unclassified); MFM (Metabolic formation); THU
     (Therapeutic use); BIOL (Biological study); FORM (Formation,
     nonpreparative); USES (Uses)
        (A; humoral and cellular immunity-enhancing agent and food)
ΙT
     Immunoglobulins
     RL: BSU (Biological study, unclassified); MFM (Metabolic formation); THU
```

(Therapeutic use); BIOL (Biological study); FORM (Formation,

```
nonpreparative); USES (Uses)
        (G; humoral and cellular immunity-enhancing agent and food)
ΙT
     Immunoglobulins
    RL: BSU (Biological study, unclassified); MFM (Metabolic formation); THU
     (Therapeutic use); BIOL (Biological study); FORM (Formation,
    nonpreparative); USES (Uses)
        (M; humoral and cellular immunity-enhancing agent and food)
IT
     Immunostimulants
        (adjuvants; humoral and cellular immunity-enhancing agent and
       food)
ΙT
     Immunity
        (cell-mediated, enhancer; humoral and cellular immunity-enhancing agent
        and food)
ΙT
    Anti-infective agents
    Antitumor agents
       Immunostimulants
    Seaweed
     Spermatochnaceae
        (humoral and cellular immunity-enhancing agent and food)
ΙT
    RL: BSU (Biological study, unclassified); MFM (Metabolic formation); THU
     (Therapeutic use); BIOL (Biological study); FORM (Formation,
    nonpreparative); USES (Uses)
        (humoral and cellular immunity-enhancing agent and food)
ΙT
        (humoral, enhancer; humoral and cellular immunity-enhancing agent and
ΙT
        (product; humoral and cellular immunity-enhancing agent and
       food)
ΙT
     Interferons
    RL: BSU (Biological study, unclassified); MFM (Metabolic formation); THU
     (Therapeutic use); BIOL (Biological study); FORM (Formation,
    nonpreparative); USES (Uses)
        (.gamma.; humoral and cellular immunity-enhancing agent and
        food)
TΤ
    9072-19-9P, Fucoidan
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
    study, unclassified); FFD (Food or feed use); PUR (Purification
     or recovery); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (humoral and cellular immunity-enhancing agent and food)
TΤ
    9072-19-9P, Fucoidan
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
    study, unclassified); FFD (Food or feed use); PUR (Purification
    or recovery); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (humoral and cellular immunity-enhancing agent and food)
                                                                           7. date
RN
     9072-19-9 HCAPLUS
CN
     Fucoidan (9CI)
                    (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L89 ANSWER 13 OF 27 HCAPLUS COPYRIGHT 2003 ACS
ΑN
    1999:291159 HCAPLUS
    131:97188
DN
     Effects of polysaccharide fucoidin on cerebrospinal fluid
TТ
     interleukin-1 and tumor necrosis factor alpha in pneumococcal
    meningitis in the rabbit
ΑU
    Granert, Carl; Raud, Johan; Waage, Anders; Lindquist, Lars
     Department of Infectious Diseases, Huddinge Hospital, Huddinge, S-141 86,
CS
```

Swed.

Infection and Immunity (1999), 67(5), 2071-2074

SO

```
CODEN: INFIBR; ISSN: 0019-9567
PB
     American Society for Microbiology
DT
     Journal
LA
     English
CC
     1-7 (Pharmacology)
     The inflammatory response in bacterial meningitis is mediated by
AB
     cytokines, such as tumor necrosis factor alpha (TNF-.alpha.) and
     interleukin-1 (IL-1), which are produced in the
     subarachnoid space by different cells, e.g., leukocytes, astrocytes, and
     microglia. The recruitment of leukocytes into the cerebrospinal fluid
     (CSF) has been shown to contribute to the neurol. damage in this disease,
     a process which could be enhanced by treatment with antibiotics.
     study, we have used a rabbit meningitis model for two sets of expts. with
     intracisternal (i.c.) injections of Streptococcus pneumoniae. First,
     pneumococcal cell wall (PCW) components were injected i.c., inducing an
     inflammatory response with pleocytosis and increased levels of CSF
     TNF-.alpha. and IL-1 at 6 and 12 h after PCW
     injection. Treatment with fucoidin, known to inhibit leukocyte
     rolling, abolished pleocytosis and inhibited the release of TNF-.alpha.
     and IL-1. In the second expt., live pneumococcal bacteria were
     injected i.c. and treatment with one dose of ampicillin (40. mg/kg of body
     wt. i.v.) was given 16 h after induction of meningitis, causing a
     sevenfold increase in CSF leukocytes over a 4-h period. CSF IL
     -1 levels at 16 h were high but did not increase further at 20 h.
     CSF TNF-.alpha. levels were high at 16 h and tended to increase at 20 h.
     Fucoidin treatment prevented the antibiotic-induced increase of
     CSF leukocytes but had no effect on the TNF-.alpha. and IL-1
     levels. Taken together, fucoidin reduced CSF TNF-.alpha. and
     IL-1 levels in acute bacterial meningitis induced by PCW fragments
     but had no effect later in the course of the disease, when live bacteria
     were used and an inflammatory increase was caused by a dose of
     antibiotics.
     polysaccharide fucoidan cerebrospinal fluid interleukin
ST
     TNFalpha; pneumococcal meningitis fucoidan CSF
     interleukin TNFalpha; inflammation fucoidan
     pneumococcal meningitis CSF leukocyte
TΤ
     Meningitis
        (bacterial; polysaccharide fucoidan effect on CSF
        interleukin-1, TNF-.alpha. and leukocyte recruitment in
        pneumococcal meningitis in rabbit)
TΤ
     Anti-inflammatory agents
     Cerebrospinal fluid
     Leukocyte
        (polysaccharide fucoidan effect on CSF interleukin
        -1, TNF-.alpha. and leukocyte recruitment in pneumococcal meningitis in
        rabbit)
ΤТ
     Interleukin 1
     Tumor necrosis factors
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (polysaccharide fucoidan effect on CSF interleukin
        -1, TNF-.alpha. and leukocyte recruitment in pneumococcal meningitis in
        rabbit)
     9072-19-9, Fucoidin
IT
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (polysaccharide fucoidan effect on CSF interleukin
        -1, TNF-.alpha. and leukocyte recruitment in pneumococcal meningitis in
        rabbit)
              THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
       25
```

RE

(1) Arfors, K; Blood 1987, V69, P338 HCAPLUS

- (2) Carlos, T; Blood 1994, P2068 HCAPLUS(3) Espevik, T; J Immunol Methods 1986, V95, P99 HCAPLUS(4) Freyer, D; Glia 1996, V16, P1 MEDLINE
- (5) Gearing, A; J Immunol Methods 1987, V99, P7 HCAPLUS
- (6) Granert, C; Clin Diagn Lab Immunol 1998, V5, P322 HCAPLUS
- (7) Granert, C; J Clin Investig 1994, V93, P929 HCAPLUS
- (8) Heumann, D; Infect Immun 1994, V62, P2715 HCAPLUS
- (9) Lindbom, L; Acta Physiol Scand 1992, V146, P415 HCAPLUS
- (10) Lindquist, L; Scand J Infect Dis 1987, V19, P263 MEDLINE
  (11) Mosmann, T; J Immunol 1986, V136, P2348 HCAPLUS
- (12) Mosmann, T; J Immunol Methods 1983, V65, P55 MEDLINE
- (13) Mustafa, M; J Infect Dis 1989, V160, P818 HCAPLUS
- (14) Pfister, H; Clin Infect Dis 1994, V19, P463 MEDLINE
- (15) Riesenfeld-Orn, I; Infect Immun 1989, V57, P1890 HCAPLUS
- (16) Saez-Llorens, X; J Clin Investig 1991, V88, P2003 HCAPLUS
- (17) Saukkonen, K; J Exp Med 1990, V171, P439 HCAPLUS
- (18) Sharief, M; J Infect Dis 1992, V166, P350 MEDLINE
- (19) Tauber, M; J Infect Dis 1988, V157, P456 MEDLINE
- (20) Tuomanen, E; J Exp Med 1989, V170, P959 MEDLINE
- (21) Tuomanen, E; J Infect Dis 1987, V155, P985 HCAPLUS
- (22) van Furth, A; Infect Immun 1996, V64, P4883 HCAPLUS
- (23) Varki, A; Proc Natl Acad Sci USA 1994, V91, P7390 HCAPLUS
- (24) Waage, A; J Exp Med 1989, V170, P1859 HCAPLUS
- (25) Zysk, G; J Neuroimmunol 1997, V73, P77 HCAPLUS
- IT 9072-19-9, Fucoidin
  - RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
    - (polysaccharide fucoidan effect on CSF interleukin
    - -1, TNF-.alpha. and leukocyte recruitment in pneumococcal meningitis in rabbit)
- RN 9072-19-9 HCAPLUS
- CN Fucoidan (9CI) (CA INDEX NAME)
- \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*
- L89 ANSWER 14 OF 27 HCAPLUS COPYRIGHT 2003 ACS
- AN 1999:87463 HCAPLUS
- DN 130:266437
- TI Development of **Fucoidan** from Kagome seaweed as novel health food materials
- AU Sakai, Takeshi; Kato, Ikunoshin
- CS Bio Operation Dept., Takara Brewing Co., Ltd., Japan
- SO New Food Industry (1998), 40(12), 1-5 CODEN: NYFIAM; ISSN: 0547-0277
- PB Shokuhin Shizai Kenkyukai
- DT Journal; General Review
- LA Japanese
- CC 17-0 (Food and Feed Chemistry)
- AB A review with 16 refs. on **Fucoidan** which is a group of polysaccharides contg. fucose, galactose, mannose, glucuronic acid, and sulfuric acid.
- ST review Fucoian polysaccharide seaweed health food
- IT Health food

Seaweed

(development of **Fucoidan** from Kagome seaweed as novel health **food** materials)

- IT Polysaccharides, biological studies
  - RL: FFD (Food or feed use); BIOL (Biological study); USES (Uses) (development of Fucoidan from Kagome seaweed as novel health food materials)
- IT 9072-19-9, Fucoidan
  - RL: FFD (Food or feed use); BIOL (Biological study); USES (Uses)

goodate

(development of Fucoidan from Kagome seaweed as novel health food materials) ΙT 9072-19-9, Fucoidan RL: FFD (Food or feed use); BIOL (Biological study); USES (Uses) (development of Fucoidan from Kagome seaweed as novel health food materials) 9072-19-9 HCAPLUS RN Fucoidan (9CI) (CA INDEX NAME) CN \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\* L89 ANSWER 15 OF 27 HCAPLUS COPYRIGHT 2003 ACS 1998:492906 HCAPLUS ΑN DN 129:215633 Anaphylaxis-induced mesenteric vascular permeability, granulocyte TI adhesion, and platelet aggregates in rat Withers, Geoffrey D.; Kubes, Paul; Ibbotson, Geoffrey; Scott, R. Brent ΑU Gastrointestinal and Immunology Research Groups and Department of CS Pediatrics, University of Calgary, Calgary, AB, T2N 4N1, Can. American Journal of Physiology (1998), 275(1, Pt. 2), H274-H284 SO CODEN: AJPHAP; ISSN: 0002-9513 PΒ American Physiological Society DT Journal English LA 15-9 (Immunochemistry) CC This study investigates the response of small venules to IgE AB -dependent, antigen-mediated mast cell activation. Intravital microscopy was utilized to visualize 25-40-.mu.m mesenteric venules, mast cell degranulation (online detection), vascular permeability changes (albumin leakage), leukocyte adhesion, and the formation of platelet aggregates in rats sensitized with 10 .mu.g of i.p. egg albumin (EA) in saline- or sham-sensitized (saline alone) rats. Sensitized rats challenged with EA (1 mg/mL superfusing mesentery), but not sensitized rats challenged with BSA or sham-sensitized rats challenged with EA, exhibited mast cell degranulation with significant time-dependent increases in vascular permeability (inhibited by diphenhydramine, salbutamol, and indomethacin), leukocyte adhesion (inhibited by Web-2086), and the formation of cellular aggregates (platelet), which were assocd. with intermittent obstruction of venular flow. Anti-platelet antibody, but not anti-neutrophil antibody or fucoidin (selectin antagonist), prevented platelet aggregate formation. Compd. 48/80-induced mast cell degranulation caused similar changes in permeability (via different mediators) and leukocyte adhesion but did not induce platelet aggregation. EA-induced platelet aggregation was not inhibited by any of the mediators tested, and platelets isolated from sensitized rats failed to aggregate in response to direct EA challenge, suggesting release of an unidentified inflammatory mediator as the factor initiating platelet aggregation. ST intestinal anaphylaxis mesenteric vessel permeability IgE; granulocyte adhesion intestinal anaphylaxis IgE; platelet aggregation intestinal anaphylaxis IgE TΨ Immunoglobulins RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence) (E; IgE-dependent, antigen-mediated mast cell activation in intestinal anaphylaxis model) IT Cell adhesion Polymorphonuclear leukocyte (IgE-dependent, antigen-mediated mast cell activation in intestinal anaphylaxis model) TT Allergens RL: ADV (Adverse effect, including toxicity); BIOL (Biological study) (IgE-dependent, antigen-mediated mast cell activation in intestinal anaphylaxis model)

ΙT Mast cell (activation; IgE-dependent, antigen-mediated mast cell activation in intestinal anaphylaxis model) IT Platelet (blood) (aggregation; IgE-dependent, antigen-mediated mast cell activation in intestinal anaphylaxis model) IT Intestine, disease Intestine, disease (anaphylaxis; IgE-dependent, antigen-mediated mast cell activation in intestinal anaphylaxis model) ΙT Anaphylaxis Anaphylaxis (intestinal; IqE-dependent, antigen-mediated mast cell activation in intestinal anaphylaxis model) TT Cell activation (mast cell; IgE-dependent, antigen-mediated mast cell activation in intestinal anaphylaxis model) IT Blood vessel (permeability; IgE-dependent, antigen-mediated mast cell activation in intestinal anaphylaxis model) ΙT Biological transport (permeation, vascular; IgE-dependent, antigen-mediated mast cell activation in intestinal anaphylaxis model) ΙT Cell aggregation (platelet; IgE-dependent, antigen-mediated mast cell activation in intestinal anaphylaxis model) RE.CNT THERE ARE 38 CITED REFERENCES AVAILABLE FOR THIS RECORD RE (1) Augustin, R; Handbook of Experimental Immunology 1979, P45.1 (2) Batchelor, J; Lancet 1975, V1, P1169 MEDLINE (3) Blockmans, D; Blood Rev 1995, V9, P143 MEDLINE (4) Butchers, P; Br J Pharmacol 1979, V67, P23 HCAPLUS (5) Cines, D; J Immunol 1986, V136, P3433 HCAPLUS (6) Coeffier, E; J Leukoc Biol 1990, V47, P234 HCAPLUS (7) Crowe, S; Gastroenterology 1992, V103, P1075 MEDLINE (8) Gaboury, J; J Immunol 1995, V154, P804 HCAPLUS (9) Heavey, D; J Immunol 1988, V140, P1953 HCAPLUS (10) Herd, C; Eur Respir J 1994, V7, P1145 HCAPLUS (11) Hogaboam, C; Gastroenterology 1993, V104, P122 HCAPLUS (12) Inagaki, N; Life Sci 1992, V51, P201 (13) Ishizaka, T; J Allergy Clin Immunol 1978, V61, P320 HCAPLUS (14) Joseph, M; Eur J Immunol 1986, V16, P306 HCAPLUS (15) Joseph, M; Nature 1983, V303, P810 HCAPLUS (16) Kanwar, S; Microcirculation 1994, V1, P175 MEDLINE (17) Kubes, P; Am J Physiol, Gastrointest Liver Physiol 1990, V259(22), PG300 (18) Kubes, P; Am J Physiol, Heart Circ Physiol 1996, V271(40), PH2438 (19) Kubes, P; J Immunol 1994, V152, P3570 HCAPLUS (20) Kurose, I; Circ Res 1994, V74, P376 MEDLINE (21) Kurose, I; Gastroenterology 1994, V107, P70 HCAPLUS (22) Kurose, I; J Clin Invest 1994, V94, P1919 HCAPLUS (23) Lagunoff, D; J Histochem Cytochem 1972, V20, P938 HCAPLUS (24) Maclouf, J; J Biol Chem 1988, V263, P174 HCAPLUS (25) Maric, M; Can J Physiol 1989, V243, P83 (26) Orr, T; Life Sci 1971, V10, P805 HCAPLUS (27) Packham, M; Can J Physiol Pharmacol 1993, V72, P278 (28) Pearce, F; Prog Med Chem 1982, V19, P60 (29) Perdue, M; Gastroenterology 1984, V86, P391 HCAPLUS (30) Pinckard, N; J Immunol 1977, V119, P2185 (31) Pothoulakis, C; Gastroenterology 1993, V105, P701 MEDLINE (32) Scott, R; Am J Physiol, Gastrointest Liver Physiol 1988, V255(18), PG505 (33) Scott, R; Am J Physiol, Gastrointest Liver Physiol 1990, V259(22), PG6 (34) Scott, R; Can J Physiol Pharmacol 1996, V74, P320 HCAPLUS (35) Shanahan, F; Int Arch Allergy Appl Immunol 1986, V80, P424 HCAPLUS

```
(36) Sugimoto, K; Int Arch Immunol 1994, V105, P195 HCAPLUS
(37) Theoharides, T; Biochem Pharmacol 1985, V34, P1389 HCAPLUS
(38) Yamamoto, H; J Clin All Immunol 1996, V91, P79
L89
     ANSWER 16 OF 27 HCAPLUS COPYRIGHT 2003 ACS
     1998:429349 HCAPLUS
ΑN
     129:174742
DN
     Properties of fucoidin extracted from kombu and its application
TI
     to food
ΑU
     Sakai, Takeshi; Kato, Ikunoshin
CS
     Takara Shuzo K.K., Japan
     Shokuhin to Kagaku (1998), 40(6), 89-93
SO
     CODEN: SHTKAY; ISSN: 0037-4105
PΒ
     Shokuhin to Kagakusha
DT
     Journal; General Review
LA
     Japanese
CC
     17-0 (Food and Feed Chemistry)
     Section cross-reference(s): 1, 18
AB
     A review with 20 refs. on fucoidin extd. from kombu as a
     food material. The prepn. of fucoidin, the phys.
     properties, the physiol. functions, e.g., cancer cell apoptosis-inducing
     effect, the application of fucoidin to food, and the
     related researches are described.
ST
     review fucoidin kombu dietary fiber
ΙT
        (functional; properties of fucoidin extd. from kombu and
        application to food)
TT
     Laminaria
        (properties of fucoidin extd. from kombu and application to
        food)
IT
     Dietary fiber
        (properties of fucoidin extd. from kombu and application to
        food as)
ΙT
     9072-19-9, Fucoidin
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); FFD (Food or feed use); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (properties of fucoidin extd. from kombu and application to
        food)
ΙT
     9072-19-9, Fucoidin
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); FFD (Food or feed use); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (properties of fucoidin extd. from kombu and application to
        food)
RN
     9072-19-9 HCAPLUS
CN
     Fucoidan (9CI) (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     ANSWER 17 OF 27 HCAPLUS COPYRIGHT 2003 ACS
L89
ΑN
     1998:402496 HCAPLUS
DN
     129:40406
     Foods containing fucoidan for taste improvement
ΤI
ΙN
     Itaya, Yoshiro
PA
     Itaya, Yoshiro, Japan
SO
     Jpn. Kokai Tokkyo Koho, 6 pp.
     CODEN: JKXXAF
DT
     Patent
LA
     Japanese
IC
     ICM A23L001-05
         A21D002-18; A23B007-10; A23G001-00; A23G003-00; A23G009-02;
     ICS
```

A23L001-20; A23L001-22; A23L001-325; C08B037-00

```
17-6 (Food and Feed Chemistry)
CC
FAN.CNT 1
                                           APPLICATION NO.
                     KIND DATE
                                                            DATE
     PATENT NO.
                           -----
                                           -----
                     ____
                                                            -----
     -----
     JP 10165114
JP 2932170
                      A2
                            19980623
                                           JP 1996-357434
                                                            19961206 <--
PI
                     B2
                            19990809
PRAI JP 1996-357434
                           19961206 <--
     Foods contain fucoidan isolated from brown algae.
     Fucoidan improves tastes of foods by controlling
     sweetness and enhancing sourness or saltiness.
ST
     food additive fucoidan brown algae
ΙT
     Bakery products
        (cakes; foods contg. fucoidan of brown algae for
        taste improvement)
ΙT
     Bread
     Brown algae (Phaeophyceae)
     Chocolate
     Condiments
     Decapterus muroadsi
       Food additives
     Soy sauce
        (foods contg. fucoidan of brown algae for taste
        improvement)
ΙT
     Vigna angularis
        (paste; foods contg. fucoidan of brown algae for
        taste improvement)
IT
     Frozen desserts
        (sherbet; foods contg. fucoidan of brown algae for
        taste improvement)
    Soups
        (stocks; foods contg. fucoidan of brown algae for
        taste improvement)
IT
     Kumquat (Fortunella)
        (sweetened; foods contg. fucoidan of brown algae
        for taste improvement)
IT
     9072-19-9, Fucoidan
     RL: FFD (Food or feed use); MOA (Modifier or additive use); BIOL
     (Biological study); USES (Uses)
        (foods contg. fucoidan of brown algae for taste
        improvement)
     9072-19-9, Fucoidan
TT
     RL: FFD (Food or feed use); MOA (Modifier or additive use); BIOL
     (Biological study); USES (Uses)
        (foods contg. fucoidan of brown algae for taste
        improvement)
     9072-19-9 HCAPLUS
RN
     Fucoidan (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L89 ANSWER 18 OF 27 HCAPLUS COPYRIGHT 2003 ACS
AN
     1998:168535 HCAPLUS
DN
     128:293900
     Histamine-induced biphasic macromolecular leakage in the microcirculation
ΤT
     of the conscious hamster: evidence for a delayed nitric
     oxide-dependent leakage
     Gimeno, G.; Carpentier, P. H.; Desquand-Billiald, S.; Hanf, R.; Finet, M.
ΑU
     Service de Pharmacologie, Laboratoire Innothera, Arcueil, 94111, Fr.
CS
SO
     British Journal of Pharmacology (1998), 123(5), 943-951
     CODEN: BJPCBM; ISSN: 0007-1188
PB
     Stockton Press
DT
     Journal
T.A
     English
```

```
CC
    15-9 (Immunochemistry)
     Section cross-reference(s): 2
    Late effects (up to 3 h) of i.v.-injected histamine on FITC-dextran
AB
    extravasation were investigated in the conscious hamster, by use of
    computer-assisted image anal. of fluorescence distribution in a
    microscopic window of dorsal skin fold prepns. This anal. allowed
    measurement of local (skin) and general (all organs) extravasations caused
    by a bolus injection of histamine (1 mg kg-1, i.v.). Histamine doses
    higher than 0.01 mg kg-1 caused biphasic local and general extravasations.
     Initial phases developed fully within 15 min (for local) and 60 min (for
     general) and were followed by late phases beginning 90 min after histamine
     injection. Although the initial and late phases of histamine-induced
     extravasations had differential apparent reactivities to the autacoid, all
     the effects of histamine on the microcirculation (1 mg kg-1) were
     inhibited by pyrilamine (1 mg kg-1, i.v.) but not by cimetidine (1 mg
    kg-1, i.v.). Pretreatment with NG-monomethyl-L-arginine (L-NMMA, 30 mg
     kg-1, i.v.) or NG-nitro-L-arginine Me ester (L-NAME, 100 mg kg-1, i.v.)
     did not affect the initial phases but did prevent the late phases of local
     and general extravasations triggered by 1 mg kg-1 histamine. The
     inhibitory effects of L-NAME were reversed by L-arginine (30 mg kg-1) but
    not by D-arginine (30 mg kg-1) according to the enantioselectivity of
    nitric oxide synthase (NOS). A late NO-mediated venular
     dilatation occurred in response to plasma histamine. A low dose of
    aminoguanidine (1 mg kg-1, i.v.), a selective inhibitor of the inducible
     isoform of NOS (iNOS), mimicked the inhibitory effects of L-NAME on the
    late phases of histamine-induced macromol. extravasations and venular
     dilatation. Pretreatment with dexamethasone (1 mg kg-1, i.v.) prevented
    both the initial and late phases of histamine-induced extravasations.
    Fucoidan (1 or 25 mg kg-1, i.v.) prevented the late phases without
     affecting initial phases, consistent with a role for leukocytes adhesion
     in the development of the late NO-mediated effects of histamine. Thus,
     i.v. injection of histamine triggers a biphasic inflammatory
     cascade via initial activation of H1 receptors which induces a late
    NO-mediated PMN-dependent extravasation process.
ST
    histamine macromol leakage microcirculation nitric oxide
ΙT
    Histamine receptors
    RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
    BIOL (Biological study); OCCU (Occurrence)
        (H1; nitric oxide in histamine-induced biphasic
       macromol. leakage in microcirculation)
ΙT
    Neutrophil
        (adhesion; nitric oxide in histamine-induced
       biphasic macromol. leakage in microcirculation)
ΙT
     Blood vessel, disease
     Blood vessel, disease
        (microvessel, injury, leakage; nitric oxide in
        histamine-induced biphasic macromol. leakage in microcirculation)
ΤТ
    Cell adhesion
        (neutrophil; nitric oxide in histamine-induced
       biphasic macromol. leakage in microcirculation)
IT
     Inflammation
        (nitric oxide in histamine-induced biphasic
       macromol. leakage in microcirculation)
IT
     125978-95-2, Nitric oxide synthase
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (inducible; nitric oxide in histamine-induced
        biphasic macromol. leakage in microcirculation)
IT
     51-45-6, Histamine, biological studies 10102-43-9,
    Nitric oxide, biological studies
     RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)
        (nitric oxide in histamine-induced biphasic
       macromol. leakage in microcirculation)
```

- RN 10102-43-9 HCAPLUS
- CN Nitrogen oxide (NO) (8CI, 9CI) (CA INDEX NAME)

N = 0

- L89 ANSWER 19 OF 27 HCAPLUS COPYRIGHT 2003 ACS
- AN 1998:122131 HCAPLUS
- DN 128:242860
- TI Characteristics of histamine-induced leukocyte rolling in the undisturbed microcirculation of the rat mesentery
- AU Yamaki, Kohji; Thorlacius, Henrik; Xie, Xun; Lindbom, Lennart; Hedqvist, Per; Raud, Johan
- CS Department of Physiology & Pharmacology, Karolinska Institutet, Stockholm, S-171 77, Swed.
- SO British Journal of Pharmacology (1998), 123(3), 390-399 CODEN: BJPCBM; ISSN: 0007-1188
- PB Stockton Press
- DT Journal
- LA English
- CC 15-9 (Immunochemistry) The main objective here was to analyze the role and mode of action of the AΒ mast cell mediator histamine in leukocyte-endothelium interactions in small venules in vivo. For this purpose, the authors used a histol. approach (combined with intravital microscopy) that allows studies of rapid mediator-induced venular leukocyte accumulation, reflecting leukocyte rolling, in the undisturbed microcirculation of the rat mesentery where rolling is normally absent. The authors first examd. the relative importance of histamine and 5-hydroxytryptamine (5-HT) in acute mast cell-dependent leukocyte recruitment. The mast cell secretagogue compd. 48/80 (i.p. for 15 min) induced a marked venular accumulation of polymorphonuclear leukocytes (PMNL) which was almost abolished by combined histamine (H1) - and histamine2 (H2) - receptor blockade. In contrast, the 5-HT-receptor antagonist methysergide was inactive in this regard. Moreover, exogenous 5-HT was less active than exogenous histamine in evoking venular PMNL accumulation (histamine response dose-dependent; 5-HT response bell shaped). Prostaglandin D2 did not cause PMNL accumulation. The venular PMNL response to exogenous histamine peaked between 15 min and 1 h, was still elevated at 2 h, and then returned to prechallenge values after 3 h. At all time points, the histamine-induced PMNL accumulation was nearly abolished by i.v. treatment with the polysaccharide fucoidin (which blocks rolling but not firm adhesion per se), suggesting that the PMNL response to histamine was due to rolling rather than firm adhesion over the entire 3 h period. At no time point did histamine trigger accumulation of mononuclear leukocytes (MNL). To examine the role of histamine-receptors in the histamine-induced PMNL accumulation (i.e. rolling), the animals were pretreated with diphenhydramine (H1-receptor antagonist), cimetidine, or ranitidine (H2-receptor antagonists). Diphenhydramine alone inhibited the venular PMNL response to histamine by 52%, while both H2-receptor antagonists were completely inactive. However, the combination of cimetidine and diphenhydramine reduced the histamine-induced PMNL rolling by 82%. Furthermore, in contrast to an H3-receptor agonist, challenge with either the H1-receptor agonist 2-thiazolylethylamine or 2 different H2-receptor agonists (impromidine, dimaprit) was sufficient to provoke venular PMNL accumulation. Treatment with the nitric oxide -synthase inhibitor L-NAME did not affect the histamine-induced PMNL

ST

IT

ΙT

IT

IT

ΙT

ΙT

ΙT

IT

TΤ

TT

IT

ΙT

TΤ

rolling. 3 H pretreatment with dexamethasone reduced the PMNL response to histamine by 73%, and flow cytometric anal. showed that the dexamethasone treatment almost completely inhibited binding of sol. P-selectin to rat isolated PMNLs. Thus, initial leukocyte recruitment after mast cell activation in the rat mesentery is critically dependent on histamine release. The cellular response to histamine was due to PMNL rolling, involved activation of both H1- and H2-receptors, and lasted for 2-3 h. Moreover, the histamine-induced PMNL rolling was not dependent on nitric oxide synthesis, but was sensitive to glucocorticoid treatment, possibly via inhibition of expression or function of leukocyte P-selectin ligand(s). histamine leukocyte rolling microcirculation mesentery Histamine receptors RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence) (H1, H1; histamine and histamine receptors role in leukocyteendothelial interactions in microcirculation of mesentery) Histamine receptors RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence) (H2; histamine and histamine receptors role in leukocyte-endothelial interactions in microcirculation of mesentery) Selectins RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence) (P-; histamine and histamine receptors role in leukocyte-endothelial interactions in microcirculation of mesentery) Leukocyte (adhesion; histamine and histamine receptors role in leukocyte-endothelial interactions in microcirculation of mesentery) Inflammation (allergic, immediate-type; histamine and histamine receptors role in leukocyte-endothelial interactions in microcirculation of mesentery) Blood vessel (endothelium; histamine and histamine receptors role in leukocyte-endothelial interactions in microcirculation of mesentery) (histamine and histamine receptors role in leukocyte-endothelial interactions in microcirculation of mesentery) Glucocorticoids RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study) (histamine and histamine receptors role in leukocyte-endothelial interactions in microcirculation of mesentery) Cell adhesion (leukocyte; histamine and histamine receptors role in leukocyte-endothelial interactions in microcirculation of mesentery) Circulation (microcirculation, mesenteric; histamine and histamine receptors role in leukocyte-endothelial interactions in microcirculation of mesentery) (microcirculation; histamine and histamine receptors role in leukocyte-endothelial interactions in microcirculation of mesentery) Leukocyte (rolling; histamine and histamine receptors role in leukocyte-endothelial interactions in microcirculation of mesentery) 50-67-9, 5-Hydroxytryptamine, biological studies 51-45-6, Histamine, biological studies RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(histamine and histamine receptors role in leukocyte-endothelial

interactions in microcirculation of mesentery)

```
L89 ANSWER 20 OF 27 HCAPLUS COPYRIGHT 2003 ACS
     1998:94790 HCAPLUS
AN
     128:204354
DN
     Foods containing fucoidan, polysaccharides, and
TΤ
     organogermanium compound for cancer immunotherapy
     Sokabe, Tsutomu
ΙN
     Sokabe, Tsutomu, Japan; Matoba, Junji
PΑ
     Jpn. Kokai Tokkyo Koho, 4 pp.
SO
     CODEN: JKXXAF
\mathsf{DT}
     Patent
LA
     Japanese
     ICM A23L001-30
TC
     ICS A23L001-30; A61K031-28; A61K031-555; A61K031-715; A61K035-80;
          A61K035-84
     18-4 (Animal Nutrition)
CC
     Section cross-reference(s): 1
FAN.CNT 1
                    KIND DATE
                                          APPLICATION NO. DATE
     PATENT NO.
                     ____
                                           ______
                                                             _____
                     A2 19980210
     JP 10033142
                                            JP 1996-225814 19960724
PRAI JP 1996-225814
                            19960724
     The title foods contain U-fucoidan (apoptosis
     inducer), D fraction (immunostimulant), .beta.-glucan (immunostimulant), (GeCH2CH2CO2H)2O3 (interferon inducer, endorphin enhancer, etc.), and
     other polysaccharides. The active ingredients show synergistic anticancer
     activity (no data).
     fucoidan polysaccharide organogermanium food
ST
     anticancer immunostimulant; D fraction beta glucan food
     anticancer; apoptosis inducer fucoidan food anticancer
     synergistic; interferon inducer endorphin enhancer organogermanium
     food; cancer immunotherapy fucoidan polysaccharide
     organogermanium
ΙT
     Food
       Immunostimulants
        (foods contg. fucoidan, polysaccharides, and
        organogermanium compd. for cancer immunotherapy)
ΙT
        (inducer; foods contg. fucoidan, polysaccharides,
        and organogermanium compd. for cancer immunotherapy)
ΙT
     Interferons
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (inducer; foods contg. fucoidan, polysaccharides,
        and organogermanium compd. for cancer immunotherapy)
     Polysaccharides, biological studies
TT
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); FFD (Food or feed use); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (mixts. contg. fucoidan and organogermanium compd.;
        foods contg. fucoidan, polysaccharides, and
        organogermanium compd. for cancer immunotherapy)
TT
     Antitumor agents
        (synergistic; foods contg. fucoidan,
        polysaccharides, and organogermanium compd. for cancer immunotherapy)
     60118-07-2, Endorphin
IT
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (enhancement of; foods contg. fucoidan,
        polysaccharides, and organogermanium compd. for cancer immunotherapy)
     9041-22-9D, .beta.-Glucan, mixts. contg. fucoidan,
ΙT
     polysaccharides, and organogermanium compd. 9072-19-9D,
     Fucoidan, mixts. contg. polysaccharides and organogermanium compd.
     179180-23-5D, mixts. contg. fucoidan and polysaccharides
```

```
RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); FFD (Food or feed use); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (foods contg. fucoidan, polysaccharides, and
        organogermanium compd. for cancer immunotherapy)
IT
     9072-19-9D, Fucoidan, mixts. contg. polysaccharides and
     organogermanium compd.
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); FFD (Food or feed use); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (foods contg. fucoidan, polysaccharides, and
        organogermanium compd. for cancer immunotherapy)
     9072-19-9 HCAPLUS
RN
     Fucoidan (9CI)
                    (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    ANSWER 21 OF 27 HCAPLUS COPYRIGHT 2003 ACS
    ·1998:13813 HCAPLUS
ΑN
     128:47606
DN
     Fucoidan-containing foods or beverages
ΤI
     Umeda, Yoshihisa; Kihara, Hiroshi; Ikai, Katsushige; Kato,
ΙN
     Ikunoshin
     Takara Shuzo Co., Ltd., Japan; Umeda, Yoshihisa; Kihara,
PA
     Hiroshi; Ikai, Katsushige; Kato, Ikunoshin
     PCT Int. Appl., 76 pp.
SO
     CODEN: PIXXD2
DT
     Patent
T.A
     Japanese
     ICM A23L001-30
TC
     ICS A61K035-80; C07H005-10
CC
     17-13 (Food and Feed Chemistry)
FAN.CNT 1
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
                                                            DATE
                           _____
                                           _____
     WO 9747208
                      A1
                            19971218
                                           WO 1997-JP1664
                                                            19970515
PΤ
           AU, BG, BR, CA, CN, CZ, HU, JP, KR, MX, NO, NZ, PL, RO, SK, US,
             VN, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
     AU 9727898
                            19980107
                                           AU 1997-27898
                      A1
                                                            19970515
     AU 711896
                            19991021
     EP 916269
                       A1
                            19990519
                                           EP 1997-922085
                                                            19970515
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
     CN 1221320
                       Α
                            19990630
                                           CN 1997-195371
                                                            19970515
     CN 1081903
                            20020403
                       В
                                         KR 1998-708652
     KR 2000010670
                       Α
                            20000225
                                                            19981028
                                           US 2001-987715
     US 2002076431
                       Α1
                            20020620
                                                            20011115
PRAI JP 1996-171666
                       Α
                            19960612
     JP 1996-318598
                       Α
                            19961115
     WO 1997-JP1664
                            19970515
                       W
                      A1
                            19981109
     US 1998-180465
     Food and beverages are prepd. that contain
AB
     fucoidan which induces apoptosis. Seaweed contg. fucoidan
     is extd. with calcium chloride or sodium carbonate, and fucoidan
     is isolated.
ST
     fucoidan food beverage apoptosis
ΙT
     Beverages
       Food
        (contg. fucoidan for apoptosis)
ΙT
     Antitumor agents
        (food contq. fucoidan as)
     9072-19-9, Fucoidan
ΙT
```

```
RL: FFD (Food or feed use); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (foods or beverages contg. apoptosis-inducing)
                                                     10043-52-4, Calcium
     497-19-8, Sodium carbonate, biological studies
IT
     chloride, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (in manuf. of foods or beverages contg.
        apoptosis-inducing fucoidan)
TΤ
     9072-19-9, Fucoidan
     RL: FFD (Food or feed use); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (foods or beverages contg. apoptosis-inducing)
RN
     9072-19-9 HCAPLUS
CN
     Fucoidan (9CI)
                    (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    ANSWER 22 OF 27 HCAPLUS COPYRIGHT 2003 ACS
ΑN
     1997:79274 HCAPLUS
DN
     126:130108
TI
     Nitric oxide decreases lung injury after intestinal
     Terada, Lance S.; Mahr, Nancy N.; Jacobson, Eugene D.
ΑU
     University of Colorado Health Sciences Center, Denver, CO, 80262, USA
CS
     Journal of Applied Physiology (1996), 81(6), 2456-2460
SO
     CODEN: JAPHEV; ISSN: 8750-7587
PB
     American Physiological Society
DT
     Journal
LA
     English
CC
     14-7 (Mammalian Pathological Biochemistry)
     Section cross-reference(s): 2
AΒ
     After injury to a primary organ, mediators are released into the
     circulation and may initiate inflammation of remote organs. We
     hypothesized that the local prodn. of nitric oxide
     (NO) may act to limit the spread of inflammation to secondarily
     targeted organs. In anesthetized rats, 30 min of intestinal ischemia
     followed by 2 h of reperfusion (I/R) did not increase lung albumin leak.
     However, after treatment with NG-nitro-L-arginine Me ester (L-NAME),
     intestinal I/R led to increased lung leak, suggesting a protective effect
     of endogenous NO. The site of action of NO appeared to be the lung and
     not the gut because 1) after treatment with L-NAME, local delivery of NO
     to the lung by inhalation abolished the increase in intestinal I/R-induced
     lung leak; 2) L-NAME had no effect on epithelial permeability
     (51Cr-labeled EDTA clearance) of reperfused small bowel; and 3) after
     treatment with L-NAME, local delivery of NO to the gut by luminal
     perfusion did not improve epithelial permeability of reperfused
     intestines. Furthermore, L-NAME increased, and inhaled NO decreased, the
     d. of lung neutrophils in rats subjected to intestinal I/R, and treatment
     with the selectin antagonist fucoidan abolished L-NAME-induced
     lung leak in rats subjected to intestinal I/R. We conclude that
     endogenous lung NO limits secondary lung injury after intestinal I/R by
     decreasing pulmonary neutrophil retention.
ST
     nitric oxide lung injury intestine ischemia
IT
     Lung, disease
        (injury; nitric oxide from neutrophils decreases
        lung injury after intestinal ischemia)
ΙT
     Intestine, disease
        (ischemia; nitric oxide from neutrophils decreases
        lung injury after intestinal ischemia)
TΤ
     Reperfusion
        (nitric oxide from neutrophils decreases lung
        injury after intestinal ischemia)
```

IT Biological transport (permeation; nitric oxide from neutrophils decreases lung injury after intestinal ischemia) Intestine, disease IT (small, ischemia; nitric oxide from neutrophils decreases lung injury after intestinal ischemia) 10102-43-9, Nitric oxide, biological studies TT RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence); PROC (Process) (nitric oxide from neutrophils decreases lung injury after intestinal ischemia) 10102-43-9, Nitric oxide, biological studies ΙT RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence); PROC (Process) (nitric oxide from neutrophils decreases lung injury after intestinal ischemia) RN 10102-43-9 HCAPLUS Nitrogen oxide (NO) (8CI, 9CI) (CA INDEX NAME) CN N = 0L89 ANSWER 23 OF 27 HCAPLUS COPYRIGHT 2003 ACS 1997:29068 HCAPLUS DN 126:73743

- AN
- Differential inhibition of polymorphonuclear leukocyte recruitment in vivo TIby dextran sulfate and fucoidan
- Van Osselaer, N.; Rampart, M.; Herman, A. G. ΑU
- Division of Pharmacology, Faculty of Medicine, University of Antwerp CS (UIA), Wilrijk, B-2610, Belg.
- Mediators of Inflammation (1996), 5(5), 346-357 SO CODEN: MNFLEF; ISSN: 0962-9351
- PΒ Rapid Science Publishers
- Journal  $\mathsf{DT}$
- English LA
- CC 15-10 (Immunochemistry)
- Section cross-reference(s): 14 The selectin-mediated rolling of leukocytes along the endothelial cells is AB a prerequisite step followed by firm adhesion and extravasation into the

inflamed tissue. This initial contact can be suppressed by sulfated polysaccharides. The authors have studied the effect of sulfated polysaccharides on the ultimate polymorphonuclear leukocyte (PMN) recruitment and plasma leakage in rabbit skin in response to intradermal injection of various inflammatory mediators. PMN infiltration evoked by various PMN chemoattractants (fMLP, C5a desArg, LTB4, and IL-8) was inhibited after i.v. injection of dextran sulfate (25 mg/kg), heparin (2.times.90 mg/kg), or fucoidan (1 mg/kg). PMN-dependent plasma leakage was equally well reduced by the different

sulfated polymers. Vascular permeability induced by histamine or thrombin acting via a PMN-independent mechanism was not reduced. Fucoidan was the only polysaccharide able to suppress IL-1-induced PMN infiltration for 60-70%. Local administration of dextran sulfate had no effect on PMN-dependent plasma leakage. Differential inhibition of PMN recruitment was detd. after injection of dextran sulfate or

fucoidan depending on the type of insult. Therefore, different adhesion pathways are utilized during PMN recruitment in vivo in response to chemoattractants and IL-1.

ST leukocyte recruitment inflammation dextran sulfate fucoidan

```
Adhesion, biological
TT
       Inflammation
     Polymorphonuclear leukocyte
        different adhesion pathways are utilized during polymorphonuclear
        leukocyte recruitment in inflammation in response to
        chemoattractants and interleukin-1)
ΙT
     Interleukin 1
       Interleukin 8
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (different adhesion pathways are utilized during polymorphonuclear
        leukocyte recruitment in inflammation in response to
        chemoattractants and interleukin-1)
TT
     Blood vessel
        (endothelium; different adhesion pathways are utilized during
        polymorphonuclear leukocyte recruitment in inflammation in
        response to chemoattractants and interleukin-1)
     Polysaccharides, biological studies
TT
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (sulfated; different adhesion pathways are utilized during
        polymorphonuclear leukocyte recruitment in inflammation in
        response to chemoattractants and interleukin-1)
     59880-97-6
                                     80295-54-1D, Complement C 5a, dearginine
TT
                  71160-24-2, LTB4
     derivs.
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (different adhesion pathways are utilized during polymorphonuclear
        leukocyte recruitment in inflammation in response to
        chemoattractants and interleukin-1)
     9005-49-6, Heparin, biological studies
                                              9042-14-2, Dextran sulfate
IT
     9072-19-9, Fucoidan
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (different adhesion pathways are utilized during polymorphonuclear
        leukocyte recruitment in inflammation in response to
        chemoattractants and interleukin-1)
TT
     9072-19-9, Fucoidan
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (different adhesion pathways are utilized during polymorphonuclear
        leukocyte recruitment in inflammation in response to
        chemoattractants and interleukin-1)
     9072-19-9 HCAPLUS
RN
     Fucoidan (9CI)
                    (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L89 ANSWER 24 OF 27 HCAPLUS COPYRIGHT 2003 ACS
AN
     1996:266153 HCAPLUS
DN
     124:332302
     An experimental study on immunoregulatory effect of fucoidan
TΤ
     Shun, Juyun; Liu, Xiaohui; Zhang, Jisheng; Zhang, Shaolun; Yang, Xiaolin;
ΑU
     Xu, Hannian
     Norman Bethune University Medical Science, Changchun, 130021, Peop. Rep.
CS
     China
     Zhongguo Haiyang Yaowu (1995), 14(3), 990-13
SO
     CODEN: ZHYAE8; ISSN: 1002-3461
PΒ
     Shandong Haiyang Yaowu Kexue Yanjiuso
DT
     Journal
LA
     Chinese
CC
     1-7 (Pharmacology)
     Fucoidan promoted splenocyte proliferation induced by mitogen
AB
```

(Con A, PHA, LPS) and IL-1 prodn. of peritoneal macrophage by bacterial lipopolysaccharide (LPS), directly stimulated murine splenocyte to produce interferon-.gamma. (IFN-. gamma.), and enhanced IL-2 prodn. of splenocyte induced by Con A, function of T-cell, B-cell, macrophage, and natural killer cell. ST immunoregulator fucoidan lymphocyte macrophage interferon IL2 IT Cell proliferation Immunomodulators Macrophage (immunoregulatory effect of fucoidan) ΙŢ Lymphocyte (B-cell, immunoregulatory effect of fucoidan) IT Lymphocyte (T-cell, immunoregulatory effect of fucoidan) IT Lymphokines and Cytokines RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (interleukin 2, immunoregulatory effect of fucoidan) ΙT Lymphocyte (natural killer cell, immunoregulatory effect of fucoidan) ΙT (splenocyte, immunoregulatory effect of fucoidan) IT Interferons RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (.gamma., immunoregulatory effect of fucoidan) IΤ 9072-19-9, Fucoidan RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (immunoregulatory effect of fucoidan) 9072-19-9, Fucoidan RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (immunoregulatory effect of fucoidan) 9072-19-9 HCAPLUS RN Fucoidan (9CI) (CA INDEX NAME) \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\* ANSWER 25 OF 27 HCAPLUS COPYRIGHT 2003 ACS L89 ΑN 1994:69152 HCAPLUS DN 120:69152 Suppression by intradermal administration of heparin of eosinophil TIaccumulation but not edema formation in inflammatory reactions in guinea-pig skin ΑU Teixeira, M. M.; Hellewell, P. G. Dep. Appl. Pharmacol., Natl. Heart Lung Inst., London, SW3 6LY, UK CS British Journal of Pharmacology (1993), 110(4), 1496-500 SO CODEN: BJPCBM; ISSN: 0007-1188 DT Journal LΑ English CC 1-8 (Pharmacology) Heparin is widely used in the treatment of thrombotic disorders and as an AB aid in surgery. Anti-inflammatory effects of heparin have also been described. In this study, the authors have investigated the effects of locally-injected heparin on the edema formation and eosinophil accumulation induced by various inflammatory stimuli in guinea-pig skin. Heparin dose-dependently suppressed the accumulation of 111In-labeled eosinophils induced by i.d. injection of zymosan-activated

plasma (ZAP). The 111In-eosinophil accumulation induced by other

fonda 0 10 / 049419 inflammatory stimuli (compd. 48/80, platelet activating factor, interleukin-8 and in a passive cutaneous anaphylaxis reaction) was also suppressed by locally-injected heparin. Edema formation in response to these same stimuli was not altered by the local injection of heparin. Fucoidin, a neg.-charged sulfated algal polymer, had no effect on the 111In-eosinophil accumulation or edema formation induced by ZAP. Nevertheless, fucoidin significantly suppressed the edema formation induced by i.d. injection of cationic protein-contq. exts. of Schistosoma mansoni larvae. Heparin also inhibited edema induced by the exts., suggesting that both fucoidin and heparin were effectively neutralizing the cationic protein of the exts. to inhibit their edema-inducing activity. Thus, heparin significantly inhibited the accumulation of 111In-eosinophils, but not edema formation, induced by various inflammatory stimuli. This, taken together with the lack of effect of fucoidin, suggests that heparin interferes with the process of eosinophil trafficking by a mechanism that does not depend on neutralization of the charge of the stimulatory mols. heparin edema eosinophil inflammatory stimulus skin Eosinophil Skin (accumulation of, inflammatory stimuli induction of, heparin suppression of, after intradermal administration) Inflammation (edema formation and eosinophil accumulation induced by, in skin, heparin effect on, after intradermal administration) (formation of, inflammatory stimuli induction of, heparin effect on, after intradermal administration) 9005-49-6, Heparin, biological studies RL: BIOL (Biological study) (edema formation and eosinophil accumulation induced by inflammatory stimuli skin response to, after intradermal administration) ANSWER 26 OF 27 HCAPLUS COPYRIGHT 2003 ACS 1993:37346 HCAPLUS 118:37346 Selective and differential binding of interleukin (IL )-1.alpha., IL-1.beta., IL-2 and IL-6 to glycosaminoglycans Ramsden, Lawrence; Rider, Christopher C. R. Holloway and Bedford New Coll., Univ. London, Egham/Surrey, UK European Journal of Immunology (1992), 22(11), 3027-31 CODEN: EJIMAF; ISSN: 0014-2980 Journal English

DT LA

ST

IΤ

TT

ΙT

ΙT

L89

ΑN DN

ΤI

ΑU

CS

SO

AΒ

CC 15-5 (Immunochemistry)

The binding of interleukin (IL)-1.alpha., IL -1.beta., IL-2 and IL-6 to acidic polysaccharides was investigated by affinity chromatog. of the recombinant, radioiodinated interleukins on columns of immobilized polysaccharide. Each interleukin showed selective binding retention. Overall, heparin bound all four interleukins significantly, whereas chondroitin sulfate provided little retention. IL-1.alpha. and IL -1.beta. showed differential binding, with only the latter binding to hyaluronic acid. IL-2 was virtually completely retained on fucoidan. Noniodinated recombinant IL-2 bound similarly to fucoidan, and fucoidan was found to sequester IL-2 activity in a bioassay employing IL-2-dependent CTLL cells. In all other cases tested, interleukin retention was partial, implying that interleukin binding sites are sparsely distributed along the polysaccharide chains. suggest that during the immune response, interleukins will tend

to be retained at sites of secretion by interaction with qlycosaminoglycans in the extracellular matrix and on cell surfaces. interleukin binding glycosaminoglycan ST Glycosaminoglycans, biological studies IT RL: BIOL (Biological study) (interleukins binding to) ΙT Molecular association (of interleukins with glycosaminoglycans) Polysaccharides, biological studies ΙT RL: BIOL (Biological study) (acidic, interleukins binding to) ΙT Lymphokines and Cytokines RL: PROC (Process) (interleukin 1.alpha., binding of, to glycosaminoglycans) ITLymphokines and Cytokines RL: PROC (Process) (interleukin 1.beta., binding of, to glycosaminoglycans) Lymphokines and Cytokines IT RL: PROC (Process) (interleukin 2, binding of, to glycosaminoglycans) IT Lymphokines and Cytokines RL: PROC (Process) (interleukin 6, binding of, to glycosaminoglycans) 9004-61-9, Hyaluronic acid 9005-49-6, Heparin, biological studies TΤ 9042-14-2, Dextran sulfate 24967-93-9, Chondroitin sulfate A RL: BIOL (Biological study) (interleukins binding to) ANSWER 27 OF 27 HCAPLUS COPYRIGHT 2003 ACS 1992:529793 HCAPLUS ΑN DN 117:129793 ΤI GMP-140 (P-selectin/CD62) binds to chronically stimulated but not resting CD4+ T lymphocytes and regulates their production of proinflammatory cytokines Damle, Nitin K.; Klussman, Kerry; Dietsch, Mary T.; Mohagheghpour, Nahid; ΑU Aruffo, Alejandro CS Bristol-Myers Squibb Pharm. Res. Inst., Seattle, WA, 98121, USA European Journal of Immunology (1992), 22(7), 1789-93 SO CODEN: EJIMAF; ISSN: 0014-2980 DTJournal LAEnglish ·CC 15-10 (Immunochemistry) AΒ GMP-140, a 140-kDa granular membrane glycoprotein localized to the .alpha. granules of platelets and the Weibel-Palade bodies of endothelial cells, is thought to play an important role in adhesive interactions predominantly between granulocytes, platelets, and vascular endothelial cells during inflammation. Although GMP-140 binds to granulocytes, its binding to lymphocytes has not been demonstrated. Using genetically engineered IgG C.gamma.1 fusion protein of the extracellular domains of GMP-140, it is demonstrated that GMP-140 binds to chronically antigen (Ag)-stimulated CD4+ T cells. Freshly isolated CD4+ T cells did not bind GMP-140, but priming and subsequent stimulation with alloantigen induced and gradually increased expression of GMP-140-reactive structures on their surface. T cells isolated from rheumatoid synovial fluids also exhibited strong binding to GMP-140. The binding of GMP-140 to primed T cells is not influenced by preactivation with phorbol 12 -myristate 13-acetate, is almost completely abolished by pretreatment of T cells with neuraminidase or trypsin, and is also strongly inhibited by EDTA, the sol. sulfated glycans dextran sulfate, fucoidan, and heparin, but not by chondroitin sulfates. In spite of its strong binding

to Ag-primed T cells, GMP-140 did not modulate the proliferative responses

of these cells to various stimuli. However, GMP-140 in conjunction with anti-T cell receptor .alpha..beta. monoclonal antibodies augmented the prodn. of granulocyte-macrophage colony-stimulating factor and inhibited the prodn. of interleukin-8 by Ag-primed T cells without influencing their tumor necrosis factor-.alpha. prodn. Thus, GMP-140 binds to chronically stimulated CD4+ T cells and differentially modulates their prodn. of proinflammatory cytokines. The ability of Ag-primed T cells to bind GMP-140 may facilitate interactions with activated platelets and endothelial cells affecting the course of inflammation.

ST GMP 140 protein T lymphocyte cytokine; P selectin lymphocyte proinflammatory cytokine

IT Inflammation

(T-cell interaction with platelets and endothelial cells in, binding of <math>GMP-140 in relation to)

IT Glycoproteins, biological studies

RL: BIOL (Biological study)

(of T-lymphocyte, as GMP-140 ligands, inflammation in relation to)

IT Lymphokines and Cytokines

RL: BIOL (Biological study)

(proinflammatory, formation of, by CD4-pos. T-cells, GMP-140 binding regulation of)

IT Glycoproteins, specific or class

RL: BIOL (Biological study)

(P-selectins, CD4-pos. T-cell formation of **proinflammatory** cytokines regulation by)

IT Lymphocyte

(T-cell, CD4-pos., proinflammatory cytokines formation by, GMP-140 regulation of)

IT Lymphokines and Cytokines

RL: FORM (Formation, nonpreparative)

(interleukin 8, formation of, by CD4-pos. T-cells,

GMP-140 binding regulation of)

IT Arthritis

(rheumatoid, synovial T-cells in human, GMP-140 binding by)

IT 83869-56-1, Granulocyte-macrophage colony-stimulating factor RL: FORM (Formation, nonpreparative)

(formation of, by CD4-pos. T-cells, GMP-140 binding regulation of)

=> fil req

FILE 'REGISTRY' ENTERED AT 09:55:49 ON 11 MAR 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the  ${\tt ZIC/VINITI}$  data file provided by InfoChem.

STRUCTURE FILE UPDATES: 10 MAR 2003 HIGHEST RN 497818-02-7 DICTIONARY FILE UPDATES: 10 MAR 2003 HIGHEST RN 497818-02-7

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting  ${\tt SmartSELECT}$  searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details:

http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> d 190 ide can

L90 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS 328081-45-4 REGISTRY RN L-Galactose, O-6-deoxy-2,4-di-O-sulfo-.alpha.-L-galactopyranosyl-CN (1.fwdarw.3)-O-6-deoxy-2,4-di-O-sulfo-.alpha.-L-galactopyranosyl-(1.fwdarw.3)-O-[6-deoxy-3-O-sulfo-.alpha.-L-galactopyranosyl-(1.fwdarw.2)]-O-6-deoxy-4-O-sulfo-.alpha.-L-galactopyranosyl-(1.fwdarw.2)-O-[6-deoxy-2,3,4-tri-O-sulfo-.alpha.-L-galactopyranosyl-(1.fwdarw.3)]-O-6-deoxy-4-Osulfo-.alpha.-L-galactopyranosyl-(1.fwdarw.3)-6-deoxy-, 2,4-bis(hydrogen sulfate) (9CI) (CA INDEX NAME) FS STEREOSEARCH C42 H72 O65 S12 MF SR CA LC STN Files: CA, CAPLUS

Absolute stereochemistry.

PAGE 2-A

[] Me

\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 134:198038

=> d his 190-

(FILE 'REGISTRY' ENTERED AT 09:53:35 ON 11 MAR 2003)

FILE 'HCAPLUS' ENTERED AT 09:53:56 ON 11 MAR 2003

FILE 'REGISTRY' ENTERED AT 09:55:26 ON 11 MAR 2003 L90 1 S 328081-45-4

FILE 'HCAPLUS' ENTERED AT 09:55:36 ON 11 MAR 2003 L91 1 S L90

FILE 'USPATFULL, USPAT2' ENTERED AT 09:55:40 ON 11 MAR 2003 L92 0 S L90

FILE 'REGISTRY' ENTERED AT 09:55:49 ON 11 MAR 2003

=> fil hcaplus FILE 'HCAPLUS' ENTERED AT 09:55:57 ON 11 MAR 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 11 Mar 2003 VOL 138 ISS 11 FILE LAST UPDATED: 10 Mar 2003 (20030310/ED)

This file contains CAS Registrý Numbers for easy and accurate substance identification.

## => d 191 all hitstr

```
L91 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2003 ACS
```

AN 2001:152496 HCAPLUS

DN 134:198038

TI Remedies containing fucoidan and/or its decomposition product

IN Tominaga, Takanari; Yamashita, Syusaku; Mizutani, Shigetoshi; Sagawa, Hiroaki; Kato, Ikunoshin

PA Takara Shuzo Co., Ltd., Japan

SO PCT Int. Appl., 73 pp. CODEN: PIXXD2

DT Patent

LA Japanese

CC 63-4 (Pharmaceuticals)

Section cross-reference(s): 17

FAN.CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE

```
WO 2001013925
                            20010301
                                           WO 2000-JP5489
                                                             20000817
PΙ
                       Α1
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
             HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU,
             LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD,
             SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU,
             ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
             CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                                             20000817
     AU 2000065934
                       Α5
                            20010319
                                          AU 2000-65934
                                                             20000817
                            20020731
                                           EP 2000-953450
     EP 1226826
                       Α1
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL
                            19990820
PRAI JP 1999-234262
                      Α
     JP 2000-69223
                            20000313
                       Α
     WO 2000-JP5489
                       W
                            20000817
     The invention relates to remedies or preventives for diseases with a need
AB
     for the regulation of the prodn. of cytokines, diseases with a need for
     the prodn. of nitrogen monoxide or allergic diseases characterized by
     contg. as the active ingredient fucoidan and/or its decompn. product; and
     foods, drinks or feeds for regulating the prodn. of cytokines, foods,
     drinks or feeds for inducing the prodn. of nitrogen monoxide, antiallergic
     foods, drinks or feeds, etc. contg. fucoidan and/or its decompn. product.
ST
     fucoidan cytokine regulation disease; antiallergy fucoidan decompn
     product; nitrogen monoxide disease fucoidan
ΙT
     Immunoglobulins
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (E, inhibitors; remedies contq. fucoidan and/or its decompn. product)
     Algae
TΤ
     Echinoderm (Echinodermata)
        (fucoidan from; remedies contq. fucoidan and/or its decompn. product)
ΙT
     Drug delivery systems
        (oral; remedies contg. fucoidan and/or its decompn. product)
ΙT
     Allergy inhibitors
     Beverages
     Feed
     Food
     Immunosuppressants
        (remedies contq. fucoidan and/or its decompn. product)
ΙT
     Cytokines
     Interferons
     Interleukin 12
     Interleukins
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (remedies contg. fucoidan and/or its decompn. product)
ΙT
     Interferons
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (.gamma.; remedies contg. fucoidan and/or its decompn. product)
ΙT
     10102-43-9, Nitrogen monoxide, biological studies
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (diseases related to prodn. of; remedies contq. fucoidan and/or its
        decompn. product)
IT
     328081-45-4P
     RL: PUR (Purification or recovery); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (remedies contg. fucoidan and/or its decompn. product)
IT
     9072-19-9, Fucoidan
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (remedies contg. fucoidan and/or its decompn. product)
```

```
THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 21
RE
(1) Dainippon Ink And Chemicals Inc; JP 09255577 A 1997 HCAPLUS
(2) Granert, C; Infect Immun 1999, V67(5), P2071 HCAPLUS
(3) Kyodo Nyugyo K K; JP 1072362 A 1998
(4) Shun, J; 1996, V14(3), P990 HCAPLUS
(5) The Australian National Universitay; JP 02502006 A
(6) The Australian National Universitay; JP 09328431 A HCAPLUS
(7) The Australian National Universitay; IL 106354 Al HCAPLUS
(8) The Australian National Universitay; CA 1316828 A1 HCAPLUS
(9) The Australian National Universitay; AT 160941 E HCAPLUS
(10) The Australian National Universitay; AT 178212 E HCAPLUS
(11) The Australian National Universitay; JP 2701904 B2 HCAPLUS
(12) The Australian National Universitay; EP 355088 A1 HCAPLUS
(13) The Australian National Universitay; EP 355088 B1 HCAPLUS
(14) The Australian National Universitay; US 5541166 A HCAPLUS
(15) The Australian National Universitay; AU 605839 B2 HCAPLUS
(16) The Australian National Universitay; EP 631784 A1 HCAPLUS
(17) The Australian National Universitay; EP 631784 B1 HCAPLUS
(18) The Australian National Universitay; IL 85145 Al HCAPLUS
(19) The Australian National Universitay; AU 8812410 A1 HCAPLUS
(20) The Australian National Universitay; WO 8805301 Al 1988 HCAPLUS
(21) Yokokawa, K; JOURNAL OF CLINICAL INVESTIGATION 1993, V92(4), P2080 HCAPLUS
ΙT
    328081-45-4P
    RL: PUR (Purification or recovery); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (remedies contg. fucoidan and/or its decompn. product)
RN
    328081-45-4 HCAPLUS
CN
    L-Galactose, O-6-deoxy-2,4-di-O-sulfo-.alpha.-L-galactopyranosyl-
    (1.fwdarw.3)-0-6-deoxy-2,4-di-0-sulfo-.alpha.-L-galactopyranosyl-
    (1.fwdarw.3)-O-[6-deoxy-3-O-sulfo-.alpha.-L-galactopyranosyl-(1.fwdarw.2)]-
    O-6-deoxy-4-O-sulfo-.alpha.-L-galactopyranosyl-(1.fwdarw.2)-O-[6-deoxy-
    2,3,4-tri-O-sulfo-.alpha.-L-galactopyranosyl-(1.fwdarw.3)]-O-6-deoxy-4-O-
    sulfo-.alpha.-L-qalactopyranosyl-(1.fwdarw.3)-6-deoxy-, 2,4-bis(hydrogen
    sulfate) (9CI) (CA INDEX NAME)
```

Absolute stereochemistry.

PAGE 2-A

[] Me

=> fil wpix FILE 'WPIX' ENTERED AT 10:08:09 ON 11 MAR 2003 COPYRIGHT (C) 2003 THOMSON DERWENT

FILE LAST UPDATED: 7 MAR 2003 <20030307/UP>
MOST RECENT DERWENT UPDATE: 200316 <200316/DW>
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

- >>> SLART (Simultaneous Left and Right Truncation) is now
   available in the /ABEX field. An additional search field
   /BIX is also provided which comprises both /BI and /ABEX <<</pre>
- >>> PATENT IMAGES AVAILABLE FOR PRINT AND DISPLAY <<<
- >>> FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES,
  SEE http://www.derwent.com/dwpi/updates/dwpicov/index.html <<<
- >>> FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE,
   PLEASE VISIT:
  http://www.stn-international.de/training center/patents/stn guide.pdf <<</pre>
- >>> FOR INFORMATION ON ALL DERWENT WORLD PATENTS INDEX USER GUIDES, PLEASE VISIT: http://www.derwent.com/userguides/dwpi\_guide.html <<<

=> d l108 all abeq tech abex tot

L108 ANSWER 1 OF 26 WPIX (C) 2003 THOMSON DERWENT

2003-058727 [05] WPIX ΑN

C2003-015211 DNC

Lyase for decomposing sulfated fucoglucuronomannan to fucoidan TΤ fraction and sulfated fucoglucuronomannan oligosaccharides useful in glycotechnology including manufacture of drugs e.g. to treat thrombosis and tumor.

DC B04 D16

IN IKAI, K; KATO, I; KIMURA, H; SAKAI, T

(TAKA-N) TAKARA HOLDINGS INC PA

CYC

PΙ WO 2002086116 A1 20021031 (200305)\* JA 67p C12N009-88

> RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZM ZW

> W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ OM PH PL PT RO RU SD SE SG SI SK SL TJ TM TN TR TT TZ UA UG US UZ VN YU ZA ZM ZW

ADT WO 2002086116 A1 WO 2002-JP3853 20020418

PRAI JP 2001-155849 20010524; JP 2001-119671 20010418

IC ICM C12N009-88

ICS C07H011-00; C12N001-20; C12P019-04

AB WO 200286116 A UPAB: 20030121

> NOVELTY - Sulfated fucoglucuronomannan oligosaccharides or their salts of formula (I) or (II) are new.

> DETAILED DESCRIPTION - Sulfated fucoglucuronomannan oligosaccharides or their salts of formula (I) or (II) are new. R = H or SO3H.

INDEPENDENT CLAIMS are also included for:

- (1) a sulfated fucoglucuronomannan lyase characterized by (a) acting on sulfated fucoglucuronomannan of brown seaweed of the Fucales order to cleave and release the alpha -D-mannosyl linkage to generate an oligosaccharide carrying an unsaturated glucuronic acid group; (b) having optimum pH at 6.5-8; and (c) having optimum temperature at 30-40 deg. C;
- (2) a process for producing the sulfated fucoglucuronomannan lyase by culturing a microorganism belonging to Fucophilus genus that can produce the enzyme before recovery from the cultured material;
- (3) a process for producing sulfated fucoglucuronomannan oligosaccharides of formula (I) or (II), or their salts, by action of the lyase on a brown seaweed of the Fucales order;
- (4) a fucoidan fraction obtained by action of the lyase on a sulfated polysaccharide mixed fraction originated from the brown seaweed prior to removing sulfated fucoglucuronomannans with reduction in the number of molecules, or collecting a fucoidan fraction;
- (5) a reagent in glycotechnology containing the sulfated fucoglucuronomannan lyase; and
- (6) sulfated fucoglucuronomannans or their salts with the physicochemical properties of (a) containing fucose, mannose and glucuronic acid as the constituting sugars; and (b) molecularly reducible by the sulfated fucoglucuronomannan lyase to form compounds of formula (I) or (II).

ACTIVITY - Cytostatic; anticoaqulant; gynecological; antiallergic; immunomodulator.

MECHANISM OF ACTION - None given.

USE - The thus produced fucoglucuronomannan oligosaccharides or fucoidan fractions are applicable in glycotechnology including manufacture of drugs to treat thrombosis, tumor, allergy or organ rejection, and to prevent chlamydia adhesion to uterine epithelial cells.

ADVANTAGE - The decomposition is highly reproducible.

Dwg.0/14 CPI

FA

AB; GI; DCN

```
MC
     CPI: B04-C02D; B04-C02X; B04-L06; B07-A02B; B12-K04; B14-F04;
          B14-G02A; B14-G02C; B14-H01; B14-N07C; B14-N14; D05-C03E;
          D05-C08; D05-H09
ABEX
     EXAMPLE - A sulfated polysaccharide mixture obtained from Fucus
     vesiculosus (brown seaweed) was treated with a crude enzyme solution of
     Fucophilus fucoidanolyticus ST-1234 strain in 25 mM imidazole
     hydrochloride buffer of pH 7 at 25 degrees C for 4 days. Sulfated
     fucoglucuronomannan oligosaccharides were isolated from the fraction with
     not more than 10,000 molecular weight by DEAE Cellufine A-800 column
     chromatography then characterized by NMR and MS.
L108 ANSWER 2 OF 26 WPIX (C) 2003 THOMSON DERWENT
     2002-619786 [67]
                       WPIX
DNC C2002-175224
ΤI
     Fucoidin ester, useful as antiviral immunoregulator, e.g. feed
     additive for fish, shrimp or pet, immunoregulator vaccine for farm animal
     or antiviral injection for farm animal and pet.
DC
     B01 C03 D13
     TANG, J; WANG, W
IN
PA
     (TANG-I) TANG J
CYC
PΙ
     CN 1344565
                   A 20020417 (200267)*
                                                     A61K047-36
ADT CN 1344565 A CN 2001-136467 20011019
PRAI CN 2001-136467
                      20011019
     ICM A61K047-36
IC
     ICS
         A61P037-02
AB
          1344565 A UPAB: 20021018
     NOVELTY - Fucoidin ester (I) is new.
          DETAILED DESCRIPTION - An INDEPENDENT CLAIM is also included for
     preparation of (I).
          ACTIVITY - Antiviral; Immunoregulator.
          No biological data available.
          MECHANISM OF ACTION - None given in source material.
          USE - (I) are used as antiviral immunoregulators, e.g. feed additive
     for fish, shrimp or pet, immunoregulator vaccine for farm animal or
     antiviral injection for farm animal and pet.
          ADVANTAGE - The technological process is simple, high yield, high
     purity and easy to realize in industrial production.
     Dwg.0/0
FS
     CPI
FΑ
     AB
MC
     CPI: B04-C02D; B04-F08; B14-A02; B14-G01; B14-G02; B14-S11;
          B14-S12; C04-F08; C14-A02; C14-F02D; C14-G01; C14-G02;
          C14-S11; C14-S12; D03-G01
TECH
                    UPTX: 20021018
     TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preparation: Preparation of (I),
     comprises:
     (i) extraction of fucoidin ester from kelp and other brown algae
     through the steps of lixiviation, centrifugal separation, reverse osmosis
     to concentrate, ethanol, precipitation, active carbon decoloring,
     ultrafiltering to desalt and refining; and
     (ii) superfine crushing, vacuum drying and other technological steps.
ABEX
     EXAMPLE - None given in source material.
L108 ANSWER 3 OF 26 WPIX (C) 2003 THOMSON DERWENT
     2002-362304 [39]
                        WPIX
DNC C2002-102541
ΤI
     Biological homeostasis maintaining agents comprise fucoidan or
     its decomposition product.
DC
     B04 D13 D21
```

```
HINO, F; KATO, I; MORIHARA, E; NISHIYAMA, E; OYASHIKI, H;
IN
     SAGAWA, H; SAKAI, T
     (TAKI) TAKARA SHUZO CO LTD
PA
CYC 96
     WO 2002022140 A1 20020321 (200239)* JA
                                              96p
PΙ
                                                     A61K031-737
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ
            NL OA PT SD SE SL SZ TR TZ UG ZW
         W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK
            DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KR KZ
            LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PH PL PT RO RU
            SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW
     AU 2001088040 A 20020326 (200251)
                                                     A61K031-737
ADT WO 2002022140 A1 WO 2001-JP7894 20010912; AU 2001088040 A AU 2001-88040
     20010912
FDT AU 2001088040 A Based on WO 200222140
PRAI JP 2001-179335 20010613; JP 2000-278712
                                                 20000913; JP 2000-295077
     20000927; JP 2000-342224
                                20001109; JP 2000-379313
                                                           20001213; JP
                   20010425
     2001-128295
IC
     ICM A61K031-737
     ICS
         A23K001-16; A23L001-29; A23L002-52; A61K035-80; A61P001-04;
          A61P001-16; A61P003-06; A61P003-08; A61P031-18; A61P035-00;
          A61P043-00; C08B037-00
     WO 200222140 A UPAB: 20020621
AΒ
     NOVELTY - Biological homeostasis maintaining agents comprise
     fucoidan, its decomposition product or their salts.
          DETAILED DESCRIPTION - Biological homeostasis maintaining agents
     comprise fucoidan, its decomposition product or their salts.
     INDEPENDENT CLAIMS are also included for:
          (i) food, drinks or feeds which maintain homeostasis and comprise
     fucoidan, its decomposition product or their salts;
          (ii) fucoidan or marine algae extract prepared by
     extracting marine algae in the presence of a reductant; and
          (iii) foods, drinks, seasonings, feeds, cosmetics and drugs
     containing the extract from (ii).
          ACTIVITY - Hepatotropic; Cardiovascular-Gen.; Antidiabetic;
     Antilipemic; Anti-HIV; Cytostatic.
          In a hydroxyproline induced liver fibrosis model in SD rats
     fucoidan was administered in drinking water at 0.5%. After 5 weeks
     amount of hydroxyproline in the liver was 336 micro g/g and liver weight
     was 14.6 g compared to 702 micro g/g and 16.5 g respectively for a control
     and 164 micro g/g and 13.2 g respectively for normal rats.
          MECHANISM OF ACTION - Hypoglycemic; Cholesterol antagonist.
          USE - As homeostasis maintaining agents for treating and preventing
     hepatic disorders (such as hepatic fibrosis), blood consistency disorders
     (e.g. for lowering blood sugar or cholesterol levels), AIDS related
     disorders and cancer.
          ADVANTAGE - Extracts are less colored and have reduced bitterness,
     lowered iodine content and fresh feel.
     Dwg.0/3
FS
     CPI
FΑ
     AB; DCN
     CPI: B04-C02D; B04-F08; B14-D02A2; B14-F09; B14-G01; B14-G01B;
MC
          B14-G02; B14-H01; B14-N12; D03-H01T2; D08-B09A1
TECH
                    UPTX: 20020621
     TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Extract: Marine algae is
     extracted in the presence of ascorbic acid or its salt, erythorbic acid or
     its salt, cysteine and/or glutathione at 30-130 degrees C for 5 minutes to
     32 hours.
ABEX
     ADMINISTRATION - Dosage is 0.01-200 mg/kg/day orally (e.g. in foods,
     drinks, feeds or drugs) or parenterally (e.g. in cosmetics).
```

EXAMPLE - Dried 'Gagomeconebu' marine algae (4 g) in calcium chloride (100

mmol/l) was extracted at 95 degrees C for 2 hours in the presence of sodium ascorbate (0.50% w/w) to give a **fucoidan** concentration of 1.45 mg/ml compared to 0.89 mg/ml in the absence of sodium ascorbate.

L108 ANSWER 4 OF 26 WPIX (C) 2003 THOMSON DERWENT

AN 2002-241900 [29] WPIX

DNC C2002-072853

TI Method for inducing, maintaining and extensively culturing antigen-specific cytotoxic T cells sustaining high-level cytotoxicity with e.g. fucoidan, useful as extremely safe cell drugs including for application in immunotherapy.

DC B04 D16

IN IDENO, M; KATO, I; SAGAWA, H

PA (TAKI) TAKARA SHUZO CO LTD

CYC 95

AB

PI WO 2002014481 Al 20020221 (200229)\* JA 99p C12N005-08

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2001078734 A 20020225 (200245) C12N005-08

ADT WO 2002014481 A1 WO 2001-JP7032 20010815; AU 2001078734 A AU 2001-78734 20010815

FDT AU 2001078734 A Based on WO 200214481

PRAI JP 2000-247072 20000816

IC ICM C12N005-08

ICS A61K035-26; A61P037-04

WO 200214481 A UPAB: 20020508

NOVELTY - Inducing cytotoxic T cells with antigen-specific cytotoxicity is by incubation of precursor cells capable of differentiation into cytotoxic T cells with antigen-presenting cells in the presence of 1 or more of compounds chosen from acidic polysaccharides, acidic oligosaccharides, acidic monosaccharides and their salts.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:

- (1) a method for maintaining cytotoxic T cells with antigen-specific cytotoxicity by continuously culturing cytotoxic T cells in the presence of 1 or more of compounds chosen from acidic polysaccharides, acidic oligosaccharides, acidic monosaccharides and their salts;
- (2) a method for extensively culturing cytotoxic T cells with antigen-specific cytotoxicity by continuously culturing cytotoxic T cells in the presence of 1 or more of compounds chosen from acidic polysaccharides, acidic oligosaccharides, acidic monosaccharides and their salts;
- (3) a method for resolving cytotoxic T cells comprising selection of cell groups containing high cytotoxic T cells with antigen-specific cytotoxicity from the culture containing cytotoxic T cells thus produced;
- (4) cytotoxic  ${\tt T}$  cells with antigen-specific cytotoxicity thus produced; and
- (5) remedies containing the cytotoxic T cells as active ingredient. ACTIVITY Immunomodulator. No biodata is given in the source material.

MECHANISM OF ACTION - Cell therapy.

USE - The method is useful for inducing, maintaining and extensively culturing antigen-specific cytotoxic T cells which are useful as extremely safe cell drugs including for application in adoptive immunotherapy.

ADVANTAGE - The method is safe, and the cultured antigen-specific cytotoxic T cells sustain a high-level cytotoxicity for safe application.  ${\rm Dwg.}\,0/1$ 

FS CPÍ

FA AB; DCN

MC CPI: B04-C02; B04-C02D; B04-C02E1; B04-C02E2; B04-C02X; B04-D01; B04-F04;

B07-A02B; B14-G03; D05-C08; D05-H01; D05-H08 UPTX: 20020508

TECH

TECHNOLOGY FOCUS - BIOLOGY - Preferred Process: During the process, extensively culturing is carried out in the presence of anti-CD3 antibody as well, particularly together with feed cells such as non-viral-infected cells.

**ABEX** 

SPECIFIC COMPOUNDS - These compounds include fucoidan, heparin, alginic acid, chondroitin sulfate A, chondroitin sulfate B, pectic acid, hyaluronic acid, fucoidan degradation products, sulfated glucose, sulfated fucose and their salts.

EXAMPLE - Peripheral blood monocytes were isolated then incubated for induction of anti-influenza memory- cytotoxic T lymphocytes (CTL) for 14 days. Cytotoxicity of the cytotoxic T cells was confirmed, and continuous culturing was also carried out.

L108 ANSWER 5 OF 26 WPIX (C) 2003 THOMSON DERWENT

2002-122201 [16] WPIX

DNC C2002-037443

ΤT Preventives or remedies for granuloma particularly due to vasoligation, containing an antagonist against a macrophage scavenger receptor e.g. antibody.

DC B04 D16

ΙN JISHAGE, K; SUZUKI, H

PA (CHUS) CHUGAI SEIYAKU KK

CYC

PΙ WO 2001095938 A1 20011220 (200216)\* JA 22p A61K045-00

> RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

> W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

> > A61K045-00

AU 2001064286 A 20011224 (200227)

ADT WO 2001095938 A1 WO 2001-JP5082 20010614; AU 2001064286 A AU 2001-64286 20010614

FDTAU 2001064286 A Based on WO 200195938

PRAI JP 2000-185942 20000616

IC ICM A61K045-00

> A61K031-711; A61K031-721; A61K031-787; A61K031-795; A61K038-38; A61K039-395; A61P015-00

AB WO 200195938 A UPAB: 20020308

> NOVELTY - Preventives or remedies for granuloma contain an antagonist against macrophage scavenger receptor as active ingredient.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:

- (1) the use of an antagonist against macrophage scavenger receptor for making preventives or remedies for granuloma; and
- (2) a method for preventing or treating granuloma by administering an antagonist against macrophage scavenger receptor to a subject.

ACTIVITY - Cytostatic.

MECHANISM OF ACTION - None given in source material.

USE - The preventives or remedies are useful for treating granuloma particularly due to vasoligation.

DESCRIPTION OF DRAWING(S) - A graph showing the chronological changes in the number of sperm in wild-type mouse and scavenger receptor knockout mouse after vasoligation. (Drawing includes non-English language text). Dwg.1/2

FS CPI

FA. AB; GI; DCN

CPI: B04-B04L; B04-C02D; B04-C03D; B04-E06; B04-G04; B04-G21; B04-N02; MC B14-H01; **B14-L06**; D05-H11A2; D05-H12D2

TECH UPTX: 20020308 TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Drugs: Granuloma particularly occurs after vasoligation. The antagonist is especially an antibody for macrophage scavenger receptor, which can be a monoclonal antibody, chimeric antibody, humanized antibody or single-stranded antibody, or is a fragment of any of the antibodies that can bind with macrophage scavenger receptor, or is an antisense nucleic acid against macrophage scavenger receptor; or is a low molecular-weight compound selected from polyvinyl sulfate, polyinosinic acid, polyxanthinylic acid, polyguanylic acid, polyG, I(1:1), dextran sulfate, fucoidin, carragheenan, bovine sulfatides, maleylated low density lipid and maleylated albumin.

ABEX

ADMINISTRATION - Administration is oral or non-oral, particularly by injection e.g. intravenous at 1-300 mg daily.

EXAMPLE - Testes of male scavenger receptor A(SR-A) knock-out mouse and wild-type mouse after vasoligation were removed for in vitro testing with use of e.g. a monoclonal antibody against 2F8:mouse SR-A, and the amount of sperm produced per testis in weight was measured and compared, with similar sperm production and inhibition of the formation of granuloma in corpus epididymis and cauda epididymis.

L108 ANSWER 6 OF 26 WPIX (C) 2003 THOMSON DERWENT

AN 2002-017727 [02] WPIX

DNC C2002-005172

TI Use of **fucoidin**, optionally in combination with an antibiotic agent, for treating arthritis, e.g. septic arthritis or rheumatoid arthritis.

DC B04

IN TARKOWSKI, A; VERDRENGH, M

PA (SAHL-N) SAHLTECH I GOETEBORG AB

CYC 22

PI WO 2001082936 A1 20011108 (200202)\* EN 31p A61K031-737 RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE TR W: CA NO US

ADT WO 2001082936 A1 WO 2001-SE962 20010504

PRAI SE 2000-1631 20000504

IC ICM A61K031-737

ICS A61P029-00

AB WO 200182936 A UPAB: 20020109

NOVELTY - The use of **fucoidin** (a sulfated fucosylated polysaccharide from seaweed), optionally in combination with an antibiotic agent, for treating arthritis is new.

ACTIVITY - Antirheumatic; antiarthritic.

MECHANISM OF ACTION - Selectin antagonist.

P-selectin deficient and control mice were injected intravenously with S.aureus CFU. Mice pretreated with fucoidin developed a significantly less severe arthritis during the first 2-3 days after bacterial inoculation. On days 2 and 3 after injection of S.aureus, the mean arthritic score for fucoidin-treated mice was 0.6 compared with 1.0 for controls. The number of mice exhibiting clinical signs of arthritis was lower in the fucoidin pretreated group during the first 4 days after injection of bacteria.

Histopathological examination of the joints confirmed clinical observations. 8 Days after injection of bacteria, 20% of **fucoidin** pretreated mice exhibited cartilage and bone destruction compared to 57% of control animals. Mean synovial hypertrophy score in the **fucoidin**-pretreated group was 2.3 compared to 3.3 in the control group. None of 6 P-selectin deficient inoculated animals exhibited any histopathological signs of arthritis 3 days after bacterial inoculation, whereas in the control group, 67% displayed synovial hypertrophy and 33% also had cartilage and bone erosion.

Immunohistochemical evaluation was carried out to determine presence

of granulocytes and macrophages in the joints of NMRI and P-selectin deficient mice administered mAb specific for L-selectin prior to bacterial inoculation. No significant differences between groups were noted, indicating that L selectin is not of major importance for extravasation of phagocytic cells into the joints in S.aureus induced arthritis.

 $\ensuremath{\mathsf{USE}}$  - For treating arthritis, particularly septic arthritis or rheumatoid arthritis (claimed).

Dwg.0/2

FS CPI

FA AB; DCN

MC CPI: B04-C02D; B14-A01; B14-C06; B14-C09; B14-L06

**ABEX** 

ADMINISTRATION - Administration is oral, rectal, by injection or by inhalation. Daily dose of **fucoidin** is preferably 15-50 mg/kg. In the treatment of septic arthritis, the daily dose of antibiotic is 1-40 mg/kg.

L108 ANSWER 7 OF 26 WPIX (C) 2003 THOMSON DERWENT

AN 2001-455916 [49] WPIX

DNC C2001-137646

TI Fucoidan-like polysaccharide composite for use as immunostimulant, antitumor agent and health food, contains dried pulverized powder of fucose and/or galactose obtained from blastema of wakame seaweed.

DC B04 D16

PA (MARU-N) MARUI BUSSAN KK

CYC 1

PI JP 2001181303 A 20010703 (200149) \* 15p C08B037-00

ADT JP 2001181303 A JP 2000-52396 20000228

PRAI JP 1999-327333 19991012

IC ICM C08B037-00

ICS A61K031-726; A61K031-737; A61K035-80; A61P037-04;

A61P043-00

AB JP2001181303 A UPAB: 20010831

NOVELTY - A **fucoidan**-like polysaccharide composite, containing fucose and/or galactose obtained by washing blastema of wakame seaweed with sea water or salt water, drying under low temperature and dry pulverizing, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) manufacturing fucoidan-like polysaccharide, which involves washing blastema of wakame seaweed with sea water or salt water, the washed blastema is dried by solar process and ground at low temperature, the ground powder is defatted, dried and extracted with water, the extract is dialyzed and dried; and
- (2) an immunostimulant comprising fucoidan-like polysaccharide as active ingredient.

ACTIVITY - Immunostimulant; cytostatic.

Natural killer cell activity of blastema of wakame seaweed was tested in BALB/c male mouse (6 weeks old). The mouse was administered with a feed containing 2 % of blastema of wakame seaweed for 10 days. The result showed that the feed containing 2 % blastema of wakame seaweed efficiently increased natural killer cell activity and suppressed reduction of immunopotentiation or biophylaxis ability. The increase in natural killer cell activity prevents mutation due to viral infection or chemicals and prevents cancer.

MECHANISM OF ACTION - Macrophage phagocytosis activity enhancer.

USE - As drug, functional foodstuff or health food for use as immunostimulant, antitumor agent as natural killer cell for self-cells which produces mutation with viral infection or chemicals, and suppresses hepatopathy.

ADVANTAGE - The fucoidan-like polysaccharide is obtained without deterioration of blastema, hence is rich in vitamins, such as

vitamin A, Bl, B2, C and niacin, minerals, such as potassium, calcium, phosphorus and iron. The contents of polysaccharide effectively maintains health by eliminating stress, restraining excitation and improving disease resistant immunity. Fucoses present in polysaccharide composite has natural killer cell activity, hence self-cell which produces mutation with viral infection or chemical is destroyed. The canceration of tissue is prevented in its early stages, hence sick prevention and heath maintenance can be potentiated. The macrophage phagocytosis ability enhancement activity of the composite enables non-specific removal of invading foreign materials. The composite enhances various specific functions of liver metabolism. The washing of blastema by sea water prevents dissolution of active constituents, thereby increasing the yield of polysaccharide. Dwg.0/18

FS CPI

FA AB; DCN

MC CPI: B04-A08; B04-A09; B04-A10; B04-C02; B10-A07; B14-G01; B14-H01; B14-H01B; D05-H13

TECH UPTX: 20010831

TECHNOLOGY FOCUS - BIOLOGY - Preferred Method: The solar dried blastema is rehydrated with water, freeze dried and pulverized. Calcium salt is added to an extract containing alginic acid, calcium alginate is collected and calcium salt is removed by adding dilute hydrochloric acid and dialyzing.

ABEX

ADMINISTRATION - Administered orally at a dose of 0.5-1000, preferably 1-300 mg/day/kg.

EXAMPLE - Blastema of wakame seaweed was cleaned in sea water and extracted. The extract was solar dried for day and night at low temperature until the moisture content was 6.2+/-0.5 %, by weight (wt.%). Subsequently freeze dried and pulverized by using stone mill to obtain powder with grain size of 35-170 mesh size. The obtained powder was added with an ethanol, water and heated at 85-90 degrees C to obtain a concentrate. The concentrate was dried to obtain a fucoidan-like polysaccharide. The composite was further added with ethanol and extracted to obtain an yield of 66 %.

L108 ANSWER 8 OF 26 WPIX (C) 2003 THOMSON DERWENT

AN 2001-432636 [46] WPIX

DNC C2001-130880

TI Cosmetics for use for hair-growth stimulants, food and drinks comprises as the active ingredients substances selected from **fucoidan**, degradation products of this, sulfated monosaccharides and salts of these. DC B03 B04 D13 D21

IN DEGUCHI, S; KATO, I; KOBAYASHI, E; MIZUTANI, S; NISHIYAMA, E; SAGAWA, H

PA (TAKI) TAKARA SHUZO CO LTD; (TAKI) TAKARA BIO INC

CYC 94

PI WO 2001039731 A1 20010607 (200146)\* JA 71p A61K007-00

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2001016486 A 20010612 (200154) A61K007-00 EP 1234568 A1 20020828 (200264) EN A61K007-00

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI TR

KR 2002067041 A 20020821 (200310) A61K007-48

ADT WO 2001039731 A1 WO 2000-JP8412 20001129; AU 2001016486 A AU 2001-16486 20001129; EP 1234568 A1 EP 2000-979012 20001129, WO 2000-JP8412 20001129; KR 2002067041 A KR 2002-706771 20020527

FDT AU 2001016486 A Based on WO 200139731; EP 1234568 A1 Based on WO 200139731

```
20001005; JP 1999-341401
PRAI JP 2000-306772
                                               19991130; JP 1999-370004
                                20000323; JP 2000-220374 20000721
     19991227; JP 2000-82738
     ICM A61K007-00; A61K007-48
IC
     ICS A61K007-06
     WO 200139731 A UPAB: 20021031
AB
     NOVELTY - Cosmetics (I) comprising substances selected from
     fucoidan, degradation products of this, sulfated monosaccharides
     and salts of these.
          ACTIVITY - Dermatological; antioxidant; endocrinal general.
          Dried fucoidan was extracted from sea-weed (konnbu). The
     obtained fucoidan ( 7g ) was dissolved in a buffer solution
     comprising sodium chloride (50 mM) and 1 % ethanol and purified. The
     obtained fucoidan was dissolved in water, and citric acid was
     added. The solution was hydrolyzed so as to obtain a degradation product
     of the fucoidan. A 3 % fucoidan ethanol solution was
     applied to male mice. The mice achieved excellent hair growth and good
     skin health.
          MECHANISM OF ACTION - None given.
          USE - (I) are for use as ingredients for lotions, emulsions, creams,
     packs, ointments, bath-shampoos, washing face or -body shampoos and hair
     shampoos, food, drinks.
          ADVANTAGE - (I) have desirable cosmetological effects on skin such as
     prevention of skin aging, improving hypersensitive skin, relieving
     itching.
     Dwg.0/1
FS
     CPI
     AB; DCN
FA
     CPI: B04-C02D; B04-C02X; B14-N17; B14-R02; D03-H01T2; D08-B03;
MC
          D08-B09A
                    UPTX: 20010815
TECH
     TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred compound : The
     fucoidan comprises a fucoidan derivative selected from
     formulae (I), (II), (III) and (IV).
     R = OH or OSO3H;
       =.
          1 or more.
     The degradation product of the fucoidan is represented by
     general formula (V), (VI) or (VII).
     Preferred Cosmetics: The hair-growth stimulants further contain minoxidil
     and/or capronium chloride. The cosmetics can be applied to e.g. food,
     drinks.
L108 ANSWER 9 OF 26 WPIX (C) 2003 THOMSON DERWENT
     2001-374426 [39]
                       WPIX
AN
DNC
    C2001-114341
TΙ
     Carbohydrate mixture for use in diabetic food compositions or
     pharmaceuticals, containing (non-)digestible base carbohydrate modified by
     coupling with other carbohydrate residue, providing delayed glucose
     release.
DC
     B05 D13 D16
ΙN
     BOEHM, G; FARWER, S; KLIEM, M; SAWATZKI, G; STAHL, B
PΑ
     (NUTR-N) NUTRICIA NV
CYC
     40
     WO 2001033973 A2 20010517 (200139)* DE
PΙ
                                              24p
                                                     A23L001-00
        RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE TR
         W: AL AU BR CA CN ID IN JP LT LV MK MX NO NZ PL RO SG SI US ZA
     DE 19954233
                  A1 20010531 (200139)
                                                     C08B037-00
     AU 2001028360 A 20010606 (200152)
                                                     A23L001-00
     EP 1229803
                  A2 20020814 (200261)
                                        DE
                                                     A23L001-00
         R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE TR
ADT
    WO 2001033973 A2 WO 2000-EP11134 20001110; DE 19954233 A1 DE 1999-19954233
     19991111; AU 2001028360 A AU 2001-28360 20001110; EP 1229803 A2 EP
     2000-993030 20001110, WO 2000-EP11134 20001110
FDT AU 2001028360 A Based on WO 200133973; EP 1229803 A2 Based on WO 200133973
```

PRAI DE 1999-19954233 19991111

IC ICM A23L001-00; C08B037-00

ICS A23L001-05; A23L001-30; A61K031-715; C08B031-00; C12P019-04

AB WO 200133973 A UPAB: 20010716

NOVELTY - A new carbohydrate (CHT) mixture (I) contains at least one modified CHT (A) or (B), obtained by coupling a conventional nutritional CHT with at least one further CHT, optionally together with at least one non-modified CHT (C).

DETAILED DESCRIPTION - A new carbohydrate (CHT) mixture (I) contains at least one modified CHT (A) or (B), obtained by coupling a conventional nutritional CHT with at least one further CHT, optionally together with at least one non-modified CHT (C). (A) comprises a digestible glucan (as digestible glucose-containing basic structure), to which at least one glucose and/or other CHT residue is coupled. (B) comprises a storage or structure CHT or its low molecular component (as non-digestible basic structure), to which at least one glucose and/or other CHT residue is coupled. The amount of glucose released from (I) in the first 90 minutes of digestion (measured in a pancreatin-based in vivo digestive system) is reduced by 10 % compared with that released from an equal weight of (C) or the non-coupled components of (A) or (B).

ACTIVITY - Antidiabetic; gastrointestinal.

MECHANISM OF ACTION - None given.

USE - The use of (I) is claimed in the nutrition of diabetics or in the production of diabetic foods or pharmaceuticals. The claims also cover a diabetic food or pharmaceutical composition containing (I) as CHT component, specifically where (I) forms 35-60 (especially 41-15) % of a liquid nutritional composition.

ADVANTAGE - (I) provides a reduced postprandial increase in blood glucose levels (due to delayed release of glucose), to give a relatively constant plasma glucose level which can be metabolized as energy source by diabetic despite their insulin deficiency. The prolonged presence of (I) in the large intestine can also stimulate the intestinal microflora and reduce digestion problems.

Dwg.0/1

FS CPI

FA AB; DCN

MC CPI: B04-C02; B10-A07; B12-M10A; B14-F09; B14-S04; **D03-H01T** TECH UPTX: 20010716

TECHNOLOGY FOCUS - FOOD - Preferred Components: The digestible base component of (A) is starch, amylose, amylopectin or dextrin (or their components); and the non-digestible base component of (B) is fructan, beta-glucan, cellulose, pectin, galacturonan, galactan, galactomannan, beta- or alpha-galacto-oligosaccharide, fucoidan, mannan, xylan, xyloglucan, laminarin, chitin, chitosan, hyaluronic acid, chondroitin, proteoglycan, glucurono-oligosaccharide, arabinan, arabinoxylan, arabinogalactan, rhamno-oligosaccharide, xanthan, alginate, agar, carrageenan, hemicellulose, vegetable gum, enzymatically prepared carbohydrate, bacterial carbohydrate, N- or O-glycoprotein oligosaccharide or glycolipid oligosaccharide. (A) or (B) is prepared by enzymatic coupling of the starting CHT's.

Preferred Modified Carbohydrate (A): (A) is a maltodextrin which:
(i) is derivatized using a transglucosidase to form glycosidic bonds with glucose in the alphal - 2, alphal - 3, alphal - 4 or alphal - 6 position in a transglycosylation reaction (especially using dextransucrase from Leuconostoc mesenteriodes (EC 2.4.1.24) as transglucosidase and sucrose (preferably in excess) as glucose source);

- (ii) is derivatized using beta-galactosidase to form glycosidic bonds with galactose in the betal 3, betal 4 or betal 6 position (especially using lactose and/or melibiose (preferably in excess) as galactose source); or
- (iii) has a CGT glucan residue transferred from amylose or amylopectin (derived from starch) onto a free hydroxy group in the C2, C3 or C4 position using cyclomaltodextrin-glucanotransferase CGT (EC 2.4.1.19) from

Bacillus macerans.

Preferred Modified Carbohydrate (B): (B) is a fructan which: (i) is derivatized using a transglucosidase to form glycosidic bonds with glucose in the alphal - 2, alphal - 3, alphal - 4 or alphal - 6 position in a transglycosylation reaction (especially using transglycosidase from

an a transglycosylation reaction (especially using transglycosidase fro Aspergillus niger (EC 2.4.1.24) and maltose (preferably in excess) as glucose source); or

(ii) has a CGT glucan residue transferred from amylose or amylopectin (derived from starch) onto a free hydroxy group in the C2, C3 or C4 position using cyclomaltodextrin-glucanotransferase CGT (EC 2.4.1.19) from Bacillus macerans.

**ABEX** 

EXAMPLE - A mixture of 20 g sucrose, 100 g malodextrins (water-soluble low molecular amylose of various chain lengths) and 500 ml 20 mM acetate buffer (pH 5.2) was incubated with 2000 U of dextransucrase from Leuconostoc mesenteriodes (EC 2.4.1.24) for 5 hours at 37 degrees C, so that at least one of the glucose residues of sucrose was transferred into the maltodextrin units. After denaturation of the enzyme for 5 minutes at 100 degrees C, the obtained glucose-modified maltodextrin was purified and recovered by ultrafiltration.

L108 ANSWER 10 OF 26 WPIX (C) 2003 THOMSON DERWENT

AN 2001-349071 [37] WPIX

DNC C2001-108349

TI Non-ulcer dyspepsia foodstuff for use as health food, such as fermented milk, confectioneries, carbonated beverages, noodles, soybean milk, soup, coffee and ice creams, contains **fucoidan** extracted from brown algae.

DC D13

PA (HONS) YAKULT HONSHA KK

CYC :

PI JP 2001095528 A 20010410 (200137)\* 8p A23L001-30

ADT JP 2001095528 A JP 1999-272232 19990927

PRAI JP 1999-272232 19990927

IC ICM A23L001-30

ICS A23L002-38; A23L002-52

ICA A61K031-715; A61K035-78; A61K035-80; A61P001-04

AB JP2001095528 A UPAB: 20010704

NOVELTY - Non-ulcer dyspepsia foodstuffs contains **fucoidan** extracted from a brown algae.

USE - As foodstuffs for non-ulcer dyspepsia symptoms (claimed). For use as health food such as fermented milk, lactic acid bacteria drink, milk drink, butter, cheese, soup, carbonated beverages, ice creams, other dairy products, fruit drink, black tea, coffee, isotonic drink, non-sugar tea, cocoa, shiruko (sweet red-bean soup), fermented rice drink, refreshing drinks, soybean milk, noodles, tofu, fresh confectionery, tablet confectionery, confectionery, iced confectionery, granule and capsule.

ADVANTAGE - The non-ulcer dyspepsia foodstuffs have no side effects and can be ingested easily. Drink which contained **fucoidan** extracted from mozuku seaweed, and extracts of hub tea, persimmon tea, Hottuynia cordata and/or fennel, was taken by 20 persons with gastric tone (non-ulcer dyspepsia symptom), 4 times a day. The gastric tone symptom was efficiently improved by the drink containing **fucoidan** extract, when compared with a placebo group. Dwg.0/1

FS CPÍ

FA AB

MC CPI: D03-H01G; D03-H01T2

TECH UPTX: 20010704

TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Foodstuff: The foodstuff further comprises plant extracts such as hub tea, persimmon leaf, Hottuynia cordata and/or fennel, preferably tea extract.

ABEX

EXAMPLE - Mozuku seaweed was added with aqueous hydrochloric acid solution (2 mole), and heated for 60 minutes, to elute **fucoidan**. The elute was centrifuged. 1M sodium hydroxide was added, and the concentrated liquid was spray-dried to obtain a powder with 75% of **fucoidan** content. 30-100 g of hub tea, persimmon tea, Hottuynia cordata and/or fennel, were added to 1 kg of ion exchange water at 90degreesC. Extraction was performed for 10 minutes, filtered, and cooled to 30degreesC, to obtain a drink for non-ulcer dyspepsia symptoms.

L108 ANSWER 11 OF 26 WPIX (C) 2003 THOMSON DERWENT

AN 2001-234976 [24] WPIX

DNC C2001-070366

TI Agents for preventing or treating diseases requiring the regulation of the production of cytokines comprise **fucoidan** or its decomposition product.

DC B05 D13

IN KATO, I; MIZUTANI, S; SAGAWA, H;
TOMINAGA, T; YAMASHITA, S

PA (TAKI) TAKARA SHUZO CO LTD

CYC 94

PI WO 2001013925 A1 20010301 (200124)\* JA 73p A61K031-737

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TZ UG ZW .

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2000065934 A 20010319 (200136) A61K031-737 EP 1226826 A1 20020731 (200257) EN A61K031-737

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI

KR 2002031411 A 20020501 (200270) A61K031-737

ADT WO 2001013925 A1 WO 2000-JP5489 20000817; AU 2000065934 A AU 2000-65934 20000817; EP 1226826 A1 EP 2000-953450 20000817, WO 2000-JP5489 20000817; KR 2002031411 A KR 2002-702220 20020220

FDT AU 2000065934 A Based on WO 200113925; EP 1226826 A1 Based on WO 200113925 PRAI JP 2000-69223 20000313; JP 1999-234262 19990820

IC ICM A61K031-737

ICS A61K035-56; A61K035-80; A61P037-02; A61P037-08;
 A61P043-00

ICA C08B037-00

ICI C08B037:00

AB WO 200113925 A UPAB: 20010502

NOVELTY - Agents for preventing or treating diseases requiring the regulation of the production of cytokines, diseases requiring the production of nitric oxide or allergic diseases comprise **fucoidan** and/or its decomposition product.

DETAILED DESCRIPTION - Agents for preventing or treating diseases requiring the regulation of the production of cytokines, diseases requiring the production of nitric oxide or allergic diseases comprise fucoidan and/or its decomposition product. INDEPENDENT CLAIMS are also included for foods, drinks or feeds that

- (1) regulate the production of cytokines;
- (2) induce the production of nitric oxide; or
- (3) have antiallergic activity comprising **fucoidan** or its decomposition product.

ACTIVITY - Antiallergic.

MECHANISM OF ACTION - Cytokine-Agonist; Cytokine-Antagonist; Nitric-Oxide-Agonist; Interferon-Agonist-Gamma; Interferon-Antagonist-Gamma; Interleukin-Agonist-12; Interferon-Antagonist-12; IgE-Antagonist.

Wistar rats administered 'gagome' fucoidan at 1% in drinking water had blood IgE antibody levels of less than 2-8 compared to

FS

FΑ

MC

CR

ΤI

DC

PA

PΙ

IC

AB

8-64 for a control. USE - For preventing or treating diseases requiring the regulation of the production of cytokines (preferably interferon- gamma or interleukin-12), diseases requiring the production of nitric oxide or allergic diseases (preferably due to IgE production). Dwg.0/13CPI AB; DCN CPI: B04-C02D; B14-G02A; B14-L01; B14-L03; B14-L06; B14-L07; D03-H01T2 TECH UPTX: 20010502 TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Agent: Fucoidan is obtained from seaweed and the cytokine is interferon or interleukin (preferably interferon-gamma or interleukin-12). **ABEX** ADMINISTRATION - Dosage is 0.01-2000 mg/kg/day orally or parenterally. L108 ANSWER 12 OF 26 WPIX (C) 2003 THOMSON DERWENT 2001-015732 [02] WPIX 2000-558250 [51] DNC C2001-004178 Agents for treating and preventing diseases by inducing growth factor production comprise e.g. acidic polysaccharide or acidic sugar alcohol. B05 D13 D21 KATO, I; KOBAYASHI, E; LI, T; MIZUTANI, S; NISHIMURA, K; NISHIYAMA, E; OHNOGI, H; SAGAWA, H; SAKAI, T; WU, H (TAKI) TAKARA SHUZO CO LTD CYC 92 WO 2000062785 A1 20001026 (200102)\* JA 158p A61K031-737 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SL SZ TZ UG ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW AU 2000038373 A 20001102 (200107) A61K031-737 EP 1175907 A1 20020130 (200216) EN A61K031-737 R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI KR 2002004997 A 20020116 (200249) A61K031-70 CN 1355703 Α 20020626 (200263) A61K031-737 JP 2000611921 X 20020723 (200263) A61K031-737 ADT WO 2000062785 A1 WO 2000-JP2432 20000414; AU 2000038373 A AU 2000-38373 20000414; EP 1175907 A1 EP 2000-917309 20000414, WO 2000-JP2432 20000414; KR 2002004997 A KR 2001-712954 20011011; CN 1355703 A CN 2000-809051 20000414; JP 2000611921 X JP 2000-611921 20000414, WO 2000-JP2432 20000414 FDT AU 2000038373 A Based on WO 200062785; EP 1175907 A1 Based on WO 200062785; JP 2000611921 X Based on WO 200062785 PRAI JP 2000-99941 20000331; JP 1999-108067 19990415; JP 1999-108499 19990415; JP 1999-114542 19990422; JP 1999-129163 19990510; JP 19990521; JP 1999-154662 1999-142343 19990602; JP 1999-200982 19990714; JP 1999-275231 19990928; JP 1999-375606 19991228 ICM A61K031-70; A61K031-737 A23K001-16; A23L001-29; A23L002-52; A61K007-40; A61K031-7016; ICS A61K031-702; **A61P043-00** C08B037-00 ICA ICI C08B037:00 WO 200062785 A UPAB: 20021001 NOVELTY - Agents for treating and preventing diseases by inducing growth factor production comprises an acidic polysaccharide or its degradation product, acidic oligosaccharide, acidic monosaccharide or an acidic sugar alcohol or their salts.

DETAILED DESCRIPTION - Agents for treating and preventing diseases by

inducing growth factor production comprises an acidic polysaccharide or its degradation product, acidic oligosaccharide, acidic monosaccharide or an acidic sugar alcohol or their salts.

INDEPENDENT CLAIMS are also included for foods, drinks, feeds and cosmetics for inducing the production of growth factor comprising an acidic polysaccharide or its degradation product, acidic oligosaccharide, acidic monosaccharide or an acidic sugar alcohol or their salts.

ACTIVITY - Hepatotropic; Antiinflammatory; Respiratory-Gen.; Nootropic; Cerebroprotective; Antidiabetic; Antiparkinsonian; Ophthalmological.

MECHANISM OF ACTION - Growth-Factor-Agonist; HGF-Agonist; NGF-Agonist; IGF-Agonist. In a cell culture sulfated starch sodium salt at 10 micro g/ml increased human hepatocyte growth factor production by MRC-5 cells by 781%.

USE - For treating and preventing diseases by inducing growth factor production (preferably hepatocyte growth factor, nerve growth factor or insulin derived growth factor) such as hepatitis, liver cirrhosis, fatty liver, nephritis, pneumonia, dementia (such as Alzheimer's disease), cerebral vascular disorders, disorders due to head injury, diabetic complications, Parkinson's disease, and retinal pigmentation disorders. Dwg.0/2

FS CPI

FA AB; DCN

MC CPI: B04-C02; B04-C02X; B04-H06; B10-A07; B14-F02D1; B14-J01A3; B14-J01A4; B14-K01; B14-L01; B14-N03; B14-N10; B14-N12; B14-S04; D03-G01; D03-H01T2; D08-B

TECH UPTX: 20010110

TECHNOLOGY FOCUS - PHARMACEUTICALS - Active Agent: Growth factor production inducer is (i) a sulfated polysaccharide (preferably of seaweed, animal, plant, microbe, fish or synthetic origin, especially fucoidan); (ii) sulfated glucose, galactose, xylose, 2-deoxyglucose, tallose or mannose; or (iii) a sulfated oligosaccharide (16 are listed in the claims e.g. sulfated maltose or sulfated dideoxymaltohexaose.

ABEX

ADMINISTRATION - Dosage is 0.01-2000 mg/kg/day orally or parenterally.

L108 ANSWER 13 OF 26 WPIX (C) 2003 THOMSON DERWENT

AN 2000-681105 [67] WPIX

DNC C2000-207282

TI Compositions to deliver compounds into cells e.g. to treat rheumatoid arthritis, comprise organic halide, targeting ligand and nuclear localization sequence in combination with compound and carrier.

DC A96 B07 D16

IN MCCREERY, T; SADEWASSER, D A; UNGER, E C

PA (IMAR-N) IMARX PHARM CORP

CYC 25

PI EP 1046394 A2 20001025 (200067)\* EN 78p A61K009-127 R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI

ADT EP 1046394 A2 EP 2000-303249 20000418

PRAI US 1999-294623 19990419

IC ICM A61K009-127

ICS A61K048-00; C12N015-88

AB EP 1046394 A UPAB: 20001223

NOVELTY - Compositions for delivering compounds into cells comprise: an organic halide; a targeting ligand; and a nuclear localization sequence in combination with the compound to be delivered.

ACTIVITY - Immunoregulatory; anti-inflammatory; anti-arthritic. USE - The compositions are used to deliver compounds into cells (claimed), particularly for the treatment of autoimmune disorders and inflammatory conditions such as rheumatoid arthritis. They may also be used to deliver pharmaceuticals, drugs, diagnostic agents, synthetic

organic molecules, peptides, proteins, vitamins, steroids, genetic materials and other bioactive agents e.g. mitotic inhibitors (vinca alkaloids), radiopharmaceuticals (radioactive iodine, phosphorus and cobalt isotopes), hormones (progestins, estrogens, anti-estrogens), anthelmintics, antimalarials, antituberculotics, biologicals (immune sera, antitoxins, antivenoms), rabies prophylactic products, bacterial vaccines, viral vaccines, aminoglycosides, respiratory products (xanthine derivatives, theophylline, aminophylline), thyroid therapeutics (iodine salts, antithyroid agents), cardiovascular products (chelating agents, mercurial diuretics, cardiac glycosides), glucagons, blood products (parenteral iron, hemin, hematoporphyrins and derivatives), targeting ligands (peptides, antibodies, antibody fragments), biological response modifiers (muramyl dipeptide, muramyl tripeptide, microbial cell wall components, lymphokines - bacterial endotoxin e.g. lipopolysaccharide and macrophage activation factor), subunits of bacteria (Mycobacteria, Comebacteria), synthetic dipeptides (N-acetyl-muramyl-L-alanyl-Disoglutamine), antifungals (ketoconazole, nystatin, griseofulvin, flucytosine, miconazole, amphotericin B), toxins (ricin), immunosuppressants (cyclosporins), antibiotics ( beta -lactam, sulfazecin), hormones (growth hormone, melanocyte-stimulating hormone, estradiol, beclomethasone dipropionate, betamethasone, betamethasone acetate, betamethasone sodium phosphate, betamethasone disodium phosphate, cortisone acetate, dexamethasone, dexamethasone acetate, dexamethasone sodium phosphate, flunisolide, hydrocortisone, hydrocortisone acetate, hydrocortisone cypionate, hydrocortisone sodium phosphate, hydrocortisone sodium succinate, methylprednisolone, methylprednisolone acetate, methylprednisolone sodium succinate, paramethasone acetate, prednisolone acetate, prednisolone sodium phosphate, prednisolone tebutate, prednisone, triamcinolone, triamcinolone acetonide, triamcinolone diacetate, triamcinolone hexacetonide, fluorocortisone acetate, oxytocin, vasopressin and their derivatives), vitamins (cyanocobalamin neionic acid), retinoids and their derivatives (retinal palmitate, alpha -tocopheryl), peptides and enzymes (manganese superoxide dismutase, alkaline phosphatases), anti-allergens (amelexanox), anticoagulants (phenprocoumon, heparin), tissue plasminogen activators, streptokinase and urokinase), circulatory drugs (propranolol), metabolic potentiators (glutathione), antibiotics (p-aminosalicylic acid, isoniazid, capreomycin sulfate, cycloserine, ethambutol hydrochloride, ethionamide, pyrazinamide, rifampicin, streptomycin sulfate dapsone, chloramphenicol, neomycin, ceflacor, cefadroxil, cephalexin, cephadrine erythromycin, clindamycin, lincomycin, amoxicillin, ampicillin, bacampicillin, carbenicillin, dicloxicillin, cyclacillin, picloxicillin, hetacillin, methicillin, nafcillin, oxacillin, penicillin (G and V), ticarcillin, rifampin, tetracycline), antivirals (acyclovir, ddI, foscarnet, zidovudine, ribavirin, vidarabine monohydrate), antianginals (diltiazem, nifedipine, verapamil, erythritol tetranitrate, isosorbide dinitrate, nitroglycerin (glyceryl trinitrate), pentaerythritol tetranitrate, anti-inflammatories (difluisal, ibuprofen, indomethacin, meclofenamate, mefenamic acid, naproxen, oxyphenbutazone, phenylbutazone, piroxicam, sulindac, tolmetin, aspirin, salicylates), antiprotozoans (chloraquine, hydroxychloraquine, metronidazole, quinine, meglumine antimonate), antirheumatics (penicillamine), narcotics (paregoric), opiates (codeine, heroin, methadone, morphine, opium), cardiac glycosides (deslanoside, digitoxin, digoxin, digitalin, digitalis), neuromuscular blockers (atracurium mesylate, gallamine triethiodide, hexafluorenium bromide, metrocurine iodide, pancurium bromide, succinylcholine chloride (suxamethionium chloride), tubocurarine chloride, vencuronium bromide), sedatives (amobarbital, amobarbital sodium, aprobarbital, butabarbital sodium, chloral hydrate, ethchlorvynol, ethinamate, flurazepam hydrochloride, glutethimide, methotrimeprazine hydrochloride, methyprylon, midazolam hydrochloride, paraldehyde, pentobarbital, pentobarbital sodium, secobarbital sodium, thiopental sodium), antineoplastics (methotrexate, fluorouracil, adriamycin, mitomycin, ansamitomycin, bleomycin, cysteine arabinoside, arabinosyl

```
adenine, mercaptopolylysine, vincristine, busulfan, chlorambucil,
    azidothymidine, melphalan (e.g. PAM, L-PAM or phenylalanine mustard),
    mercaptopurine, mitotane, procarbazine hydrochloride, dactinomycin
     (actinomycin D), daunorubicin hydrochloride, dosorubicin hydrochloride,
    Taxol (RTM: paclitaxel), plicamycin (mithramycin), aminoglutethimide,
    estramustine phosphate sodium, flutamide, leuprolide acetate, megestrol
    acetate, tamoxifen citrate, testolactone, trilostane, amsacrine (m-AMSA),
    asparaginase, etoposide (VP-16), interferon alpha -2a, interferon alpha
    -2b, teniposide (VM-26), vinblastine sulfate (VLB), vincristine sulfate,
    hydroxyurea, procarbaxine or dacarbazine).
         ADVANTAGE - The compositions provide improved delivery of
    compositions including drugs and genetic materials into cells. They
    provide for specific targeting and delivery of compounds to particular
    cells and increased targeting to the nuclei of targeted cells. They also
    allow delivery to cell lines that would be otherwise resistant to
    intracellular delivery and gene expression using other conventional means.
          DESCRIPTION OF DRAWING(S) - Schematic representation of a targeted
    composition.
          targeted composition 1
    lipid coating 2
    lipids 2A
         halocarbon gas or liquid 3
          genetic material 4
          targeting ligand 5
          lipid head group 6
    tether 7
    tether 7A
         nuclear localization sequence 8
          condensing agent. 9
    Dwg.2/2
    CPI
    AB; GI; DCN
    CPI: A12-V01; B04-B04D; B04-E02D; B04-E06; B04-E07; B04-G01; B04-H01;
          B04-J01; B04-K01V; B14-A01; B14-A02; B14-A03; B14-A04; B14-B03;
          B14-C03; B14-C09B; B14-F01; B14-G02A; B14-G02D; B14-L01;
          B14-S11; D05-C10; D05-C12; D05-H12B; D05-H12D2; D05-H12D4; D05-H12D5
TECH
                    UPTX: 20001223
    TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred organic halide - The
    organic halide is a gaseous or liquid organic halide, preferably a liquid
    or a gaseous precursor. The organic halide is a fluorinated compound,
    preferably a perfluorinated compound, more preferably a perfluorocarbon,
    especially a perfluoroether compound. The organic halide is
     1-bromo-nonafluorobutane, perfluorooctyliodide, perfluorooctylbromide,
     1-chloro-1-fluoro-1-bromomethane, 1,1,1-trichloro-2,2,2-trifluoroethane,
    1,2-dichloro-2,2-difluoroethane, 1,1-dichloro-1,2-difluoroethane,
     1,2-dichloro-1,1,3-trifluoropropane, 1-bromoperfluorobutane,
     1-bromo-2, 4-difluorobenzene, 2-iodo-1, 1, 1-trifluoroethane,
     5-bromovalerylchloride, 1,3-dichlorotetrafluoroacetone, bromine
    pentafluoride, 1-bromo-1,1,2,3,3,3-hexafluoropropane, 2-chloro-1,1,1,4,4,4-
    hexafluoro-2-butene, 2-chloropentafluoro-1,3-butadiene,
     iodotrifluoroethylene, 1,1,2-trifluoro-2-chloroethane,
     1,2-difluorochloroethane, 1,1-difluoro-2-chloroethane, 1,1-dichlorodifluor
     omethane, dibromofluoromethane, chloropentafluoroethane,
    bromochlorodifluoromethane, dichloro-1,1,2,2-tetrafluoroethane,
     1,1,1,3,3-pentafluoropentane, perfluorotributylamine,
    perfluorotripropylamine, 3-fluorobenzaldehyde, 2-fluoro-5-nitrotoluene,
     3-fluorostyrene, 3,5-difluoroaniline, 2,2,2-trifluoroethylacrylate,
     3-(trifluoromethoxy)-acetophenone, 1,2,2,3,3,4,4-octafluorobutane,
     1,1,1,3,3-pentafluorobutane, 1-fluorobutane, 1,1,2,2,3,3,4,4-
     octafluorobutane, 1,1,1,3,3-pentafluorobutane, perfluoro-4-
    methylquinolizidine, perfluoro-N-methyl-decahydroquinone,
     perfluoro-N-methyl-decahydroisoquinone, perfluoro-N-cyclohexyl-
```

pyrrolidine, perfluoroheptane, perfluorocyclohexane, perfluoromethane

FS

FΑ MC

(preferred), perfluoroethane (preferred), perfluoropropane (preferred), perfluorobutane (preferred), perfluoropentane (preferred), perfluorohexane (preferred), perfluoroheptane (preferred), perfluorooctane (preferred), perfluorononane (preferred), perfluorodecane (preferred), perfluorododecane (preferred), perfluoro-2-methyl-2-pentene (preferred), perfluorocyclohexane (preferred), perfluorodecalin (preferred), perfluorododecalin (preferred), perfluoropropylene, perfluorocyclobutane, perfluoro-2-butyne, perfluoro-2-butene, perfluorobuta-1,3-diene, perfluorobutylethyl ether (preferred), bis(perfluoroisopropyl)ether (preferred), bis(perfluoropropyl)ether (preferred), perfluorotetrahydropyran (preferred), perfluoromethyl tetrahydrofuran (preferred), perfluoro-tertiary butyl-methyl ether (preferred), perfluoro-isobutyl-methyl ether (preferred), perfluoro-n-butyl-methyl ether, perfluoro-isopropyl-methyl ether (preferred), perfluoro-n-propylmethyl ether (preferred), perfluorodiethyl ether (preferred), perfluorocyclopropyl methyl ether (preferred), perfluoromethyl ethyl ether (preferred), perfluorodimethyl ether (preferred), sulfur hexafluoride or selenium hexafluoride. TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred compositions - The compositions further comprises a carrier such as a polymer, lipid, protein or metal ion. The carrier preferably comprises a lipid, more preferably a cationic lipid, especially N-(1-(2,3-dioleoyloxy)propyl)-N,N,Ntrimethylammonium chloride. The carrier preferably comprises a polymer, more preferably a polyethylene, polyoxyethylene, polypropylene, pluronic acid or alcohol, polyvinyl, polyvinylpyrrolidone, arabinan, fructan, fucan, galactan, galacturonan, glucan, mannan, xylan, levan, fucoidan, carrageenan, galactocarolose, pectin, pectic acid, amylose, pullulan, glycogen, amylopectin, cellulose, carboxylmethylcellulose, hydroxypropylmethylcellulose, dextran, pustulan, chitin, agarose, keratan, chondroitin, dermatan, hyaluronic acid, alginic acid, homopolymer or heteropolymer containing one or more of an aldose, ketose, acid, amine, erythrose, threose, ribose, arabinose, xylose, lyxose, allose, altrose, glucose, mannose, gulose, idose, galactose, talose, erythrulose, ribulose, xylulose, psicose, fructose, sorbose, tagatose, glucuronic acid, gluconic acid, glucaric acid, galacturonic acid, mannuronic acid, guluronic acid, glucosamine, galactosamine or neuraminic acid. The carrier is Lipofectin, Lipofectamine, Transfectace, Transfectam, Cytofectin, dimyristoyloxypropyl-3dimethylhydroxyethylammonium bromide (DMRIE), dilauryloxypropyl-3dimethylhydroxyethylammonium bromide (DLRIE), GAP-DLRIE, 1,2-dioleoyloxy-3-(trimethylammonio)propane (DOTAP), dioleoylphosphatidylethanolamine (DOPE), DMEAP, DODMP, dioleoylphosphtadiylcholine (DOPC), DDAB, 2,3-dioleoyloxy-N-(2-(sperminecarboxamidoethyl)-N, N-dimethyl-1-propanaminium trifluoroacetate (DOSPA), EDLPC, EDMPC, DPH, TMADPH, cetyltrimethylammonium bromide (CTAB), lysyl-PE, 3, beta-(N, (N', N'-dimethylaminoethane) carbamoyl) cholesterol (DC-Chol), alanyl cholesterol, DCGS, dipalmitoylphosphatidylethanolamine-5carboxyspermylamide (DPPES), dicaproylphosphatidylethanolamine (DC PE), 4-dimethylaminopyridine (DMAP), dimyristoylphosphatidylethanolamine (DMPE), dioctadecylamidoglycol spermidine (DOGS), DOFIME, dipalmiotylethylphosphatidylcholine (DPEPC), Pluronic (RTM: polyethylene glycol), Tween (RTM: polysorbate), Brij (RTM: polyoxyethylene glycol), plasmalogen, phosphatidylethanolamine, phosphatidylcholine, glycerol-3-ethylphosphatidylcholine, dimethylammonium propane, trimethylammonium propane, dimethyldioctadecylammonium bromide, sphingolipids, sphingomyelin, lysolipid, glycolipid, sulfatide, glycosphingolipid, cholesterol, cholesterol ester, cholesterol salt, oil, 1,2-dioleoyl-sn-glycerol, N-succinyldioleoylphosphatidylethanolamine, 1,3-dipalmitoyl-2-succinyl-glycerol, 1,2-dipalmitoyl-sn-3succinylglycerol, palmitoylhomocysteine, 1-hexadecyl-2palmitoylglycerophosphatidylethanolamine, N, N''bis(dodecylaminocarbonylmethylene)-N, N'-bis((N, N, Ntrimethylammoniumethylaminocarbonylethylene)ethylene diamine tetraiodide,

N, N''-bis(hexad ecylaminocarbonylmethylene)-N, N, N''-tris-N, N, Ntrimethylammoniumethylaminocarbonylmethylenediethylenetriamine hexaiodide, N, N'-bis (dodecylaminocarbonylmethylene) -N, N''-bis ((N, N, Ntrimethylammoniumethylaminocarbonyl-methylene)-cyclohexylene-1,4-diaminetetraiodide, 1,1,7,7-tetra((N,N,N-tetramethylammoniumethylaminocarbonylmet hylene)-3-hexadecylaminocarbonylmethylene-1,3,7-triaazaheptane heptaiodide or N,N,N',N'-tetra-((N,N,N-trimethylammoniumethylaminocarbonylmethylene)-N'-(1,2-dioleoylglycero-3-phosphoethanolaminocarbonylmethylene) diethylene triamine tetraiodide. The carrier comprises a dioleoylphosphatidylethanolamine, fatty acid, lysolipid, phosphatidylcholine, phosphatidylethanolamine, phosphatidylserine, phosphatidylglycerol, phosphatidylinositol, sphingolipid, glycolipid, glucolipid, sulfatide, glycosphingolipid, phosphatidic acid, palmitic acid, stearic acid, arachidonic acid, oleic acid, lipid bearing a polymer, lipid bearing a sulfonated saccharide, cholesterol, tocopherol hemisuccinate, lipid with an ether-linked fatty acid, lipid with an ester-linked fatty acid, polymerized lipid, diacetyl phosphate, stearylamine, cardiolipin, phospholipid with a fatty acid of 6-8C, phospholipid with asymmetric acyl chains, 6-(5-cholesten-3b-yloxy)-1-thiob-D-galactopyranoside, digalactosyldiglyceride, 6-(5-cholesten-3betayloxy) hexyl-6-amino-6-deoxy-1-thio-b-D-galactopyranoside, 6-(5-cholesten-3b)-yloxy)hexyl-6-amino-6-deoxyl-1-thio-alpha-Dmannopyranoside, 12-(((7'-diethylamino-coumarin-3yl)carbonyl)methylamino)octadecanoic acid, N-(12-(((7'-diethylaminocoumarin-3-yl)carbonyl)methylamino)octadecanoyl)-2-aminopalmitic acid, cholesteryl (4'-trimethyl-ammonio)butanoate, Nsuccinyldioleoylphosphatidylethanolamine, 1,2-dioleoyl-sn-glycerol, 1,2-dipalmitoyl-sn-3-succinyl-glycerol, 1,3-dipalmitoyl-2succinylglycerol, 1-hexadecyl-2-palmitoylglycerophosphoethanolamine and/or palmitoylhomocysteine. The carrier comprises a phosphatidylcholine, preferably dioleoylphosphatidylcholi ne, dimyristoylphosphatidylcholine, dipentadecanoylphosphatidylcholine, dilauroylphosphatidylcholine, dipalmitoylphosphatidylcholine or distearoylphosphatidylcholine. The carrier comprises phosphatidylethanolamine, preferably dioleoylphosphatidylethanolamine. The carrier comprises a glycolipid, preferably ganglioside GM1 or GM2. The carrier comprises a lipid bearing a polymer, preferably polyethylene glycol, chitin, hyaluronic acid or polyvinylpyrrolidone, more preferably polyethylene glycol, especially a polyethylene glycol with a molecular weight of 2,000, 5,000 or 8,000. The carrier comprises a phospholipid with asymmetric acyl chains with one acyl chain of about 6 C in length and another of about 12 C in length. The carrier comprises about 82 mole % dipalmitoylphosphatidylcholine, about 8 mole % dipalmitoylphosphatidylethanolamine-polyethylene glycol 5,000 and about 10 mole % dipalmitoylphosphatidic acid. The carrier comprises a surfactant, preferably a fluorosurfactant. The compositions further comprise a telomerase. The compositions further comprise a fusion peptide. Preferred delivery compound - The compound to be delivered is a pharmaceutical agent, synthetic organic molecule, protein, peptide or genetic material, preferably a mutant gene that encodes a defective receptor chosen from tumor necrosis factor (TNF), gamma interferon (IFN gamma) or interleukin-1 (IL-1), antisense oligonucleotide (that preferably hybridizes to a nucleic acid molecule encoding a protein selected from TNF receptor, IFN gamma receptor or IL-1 receptor) or a ribozyme (a ribozyme that disrupts nucleic acid molecules encoding a protein chosen from TNF receptor, IFN gamma receptor or IL-1 receptor). Preferred targeting ligand - The targeting ligand is a protein, antibody (fragment), hormone (analog), glycoprotein, lectin, (poly)peptide, amino acid, sugar, saccharide, carbohydrate, vitamin, steroid (analog), cofactor, bioactive agent or genetic material, preferably Sialyl Lewis X (preferred), mucin, hyaluronic acid, LFA-1, VLA-4, fibrinogen, von Willebrand factor, vitronectin, VCAM-1, CD49d/CD29, methyl-alpha-Dmannopyranoside, N-formal peptide, C5a, leukotriene B4, platelet-activating factor, IL-8/NAP-1, CTAP-III, beta-thromboglobulin,

NAP-2, gro/MGSA, ENA-78, MCP-1, MAP-1alpha, beta, RANTES or I-309. Preferred nuclear localization sequence - The nuclear localization sequence is a peptide, protein, receptor, transcription factor or an enzyme, especially influenza virus nucleoprotein, karyophenin betal, human statl gene, m-importin, mouse homolog of nuclear pore targeting complex, hepatitis B virus (HBV) polymerase, glucocorticoids receptor (GlucR), interferon-regulated factors ISGF-3 and GAF, yeast mating switch/HO endonuclease promoter SW15, Drosophila melanogaster morphogen dorsal, nuclear factors NF-kappa and NF-AT, T-ag, c-rel, lamin B2, GrH receptor, c-fos, cofilin, rNFIL-6, NF-ATplc, PICA C-subunit, p42mapk/p44erk1, p90rsk, PKC-alpha, lodestar, v-jun, cyclin B (B-type cyclins), adenovirus 5 Ela protein, xnf7, PwA33, Rb-1, p53, c-myc, PTF1, HMG1/2 and tegument protein pp65 (UL83) of human cytomegalovirus. The nuclear localization sequence is a peptide comprising a defined amino acid sequence.

ABEX

SPECIFIC SEQUENCES - A total of 24 nuclear localization sequences are claimed and all are given in the specification. E.g. Pro-Lys-Lys-Arg-Lys-Val and Asn-Lys-Ile-Pro-Ile-Lys-Asp.

ADMINISTRATION - Administration may be in combination with ultrasound to the cells (claimed).

L108 ANSWER 14 OF 26 WPIX (C) 2003 THOMSON DERWENT

AN 2000-195168 [17] WPIX

DNC C2000-060469

TI Treating a condition associated with glycosaminoglycan associated molecular interaction by administration of ionic compounds.

DC B05

IN GERVAIS, F; GREEN, A M; KISILEVSKY, R

PA (NEUR-N) NEUROCHEM INC; (TOOH) UNIV QUEENS KINGSTON

CYC 87

PI WO 2000006133 A2 20000210 (200017)\* EN 108p A61K031-00 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL

OA PT SD SE SL SZ UG ZW
W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB
GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU
LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR
TT UA UG UZ VN YU ZA ZW

AU 9951894 A 20000221 (200029) A61K031-00 EP 1100487 A2 20010523 (200130) EN A61K031-185

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI

US 6310073 B1 20011030 (200172) A61K031-407 US 2002193395 A1 20021219 (200303) A61K031-44

ADT WO 2000006133 A2 WO 1999-IB1473 19990728; AU 9951894 A AU 1999-51894 19990728; EP 1100487 A2 EP 1999-936931 19990728, WO 1999-IB1473 19990728; US 6310073 B1 Provisional US 1998-94454P 19980728, US 1999-362505 19990727; US 2002193395 A1 Provisional US 1998-94454P 19980728, Cont of US 1999-362505 19990727, US 2001-970148 20011002

FDT AU 9951894 A Based on WO 200006133; EP 1100487 A2 Based on WO 200006133; US 2002193395 A1 Cont of US 6310073

PRAI US 1999-362505 19990727; US 1998-94454P 19980728; US 2001-970148 20011002

IC ICM A61K031-00; A61K031-185; A61K031-407; A61K031-44 ICS A01N043-42; A61K031-435; A61K031-47; A61K031-715; A61K038-02

AB WO 200006133 A UPAB: 20000405 NOVELTY - Treating a condition associated with a glycosaminoglycan associated molecular interaction comprises administering an ionic compound (I).

DETAILED DESCRIPTION - Treating a condition associated with a glycosaminoglycan associated molecular interaction (GAMI) comprises administering an ionic compound of formula Q(Y-X+)n (I) or its salt or ester.

Y = anionic group at physiological pH;

FS

FΑ

MC

ΑN

TΤ

DC

PΑ

PΙ

IC

AB

```
Q = a carrier molecule;
          X = cationic group; and
          n = integer selected such that the biodistribution of the
     therapeutic compound for an intended target site is not prevented while
     maintaining activity of the therapeutic compound.
          INDEPENDENT CLAIMS are also included for:
          (A) a method of modulating interaction between an infectious agent
     and a glycosaminoglycan comprising administering a therapeutic agent
     comprising at least one sulfonate group attached to a carrier molecule or
     its salt or ester (i); and
          (B) a packaged pharmaceutical composition for treating a GAMI
     comprising a container containing (i) and instructions for use.
          ACTIVITY - Antibacterial; Virucide
          MECHANISM OF ACTION - Glycosaminoglycan-Antagonist.
          USE - For treating conditions associated with a glycosaminoglycan
     associated molecular interaction such as bacterial infection
     e.g. (Chlamydia trachomatis, Staphylococcus aureus, Pseudomonas aeruginosa,
     Legionella pneumophila, Bordetella pertussis, and Mycoplasma pneumoniae
     and viral infection (e.g. infection associated with Herpes viridae such as
     herpes simplex and cytomegalovirus).
     Dwg.0/28
     CPI
     AB; DCN
     CPI: B04-C01; B04-C02; B04-C03; B05-B01G; B06-H; B07-H; B10-A09B; B11-C06;
          B14-A01; B14-A02; B14-L06
TECH
                    UPTX: 20000405
     TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Molecule: The carrier
     molecule is a carbohydrate, polymer, peptide, peptide derivative,
     aliphatic group, alicyclic group, heterocyclic group and/or aromatic group
     (preferably an aliphatic group).
     Preferred Method: GAMI does not include amyloidosis and the interaction
     with a cell surface. GAMI is associated with a bacterial or viral
     infection, provided that when the bacterium is Chlamydia trachomatis then
     (I) is not carrageenan, pentosan polysulfate, fucoidan, dextran
     sulfate, heparin, heparan sulfate or dermatan sulfate and when the viral
     infection is cytomegalovirus then (I) is not a chondroitin sulfate.
ABEX
     SPECIFIC COMPOUNDS - The use of 18 compounds is specifically claimed e.g.
     3-amino-1-propanesulfonic acid.
     ADMINISTRATION - Dosage is 5-500 mg/kg/day orally, or e.g. by injection.
L108 ANSWER 15 OF 26 WPIX (C) 2003 THOMSON DERWENT
     1999-543942 [46]
                       WPIX
DNC C1999-158979
     Immuno-potentiating agent to reinforce humoral and cellular immunity for
     prevention and treatment of infection - contains fucoidan as
     effective ingredient.
     B04 D13
     (HONS) YAKULT HONSHA KK
CYC
    1
                  A 19990824 (199946)*
                                                     C08B037-00
     JP 11228602
                                             4p
ADT JP 11228602 A JP 1998-41043 19980209
                      19980209
PRAI JP 1998-41043
     ICM C08B037-00
         A23L001-30; A61K031-725
     ICS
ICA A61K035-80
     JP 11228602 A UPAB: 19991116
     NOVELTY - Immunopotentiating agent contains fucoidan as
     effective ingredient.
          DETAILED DESCRIPTION - An INDEPENDENT CLAIM is also included for
     immunoactivity enriched food containing fucoidan.
```

USE - For prevention and treatment of infection. ACTIVITY - Immunostimulant. Mouse agmen peyerianum cells were cultivated and fucoidan was added and allowed to stand for 7 days. The supernatent liquid was removed and the IgA, IgM and IgG present was determined using ELISA. In mu g/ml, 1000 fucoidan produced 1.8 IgA, 9.05 IgM and 1.22 IgG antibodies. ADVANTAGE - The agent reinforces both humoral and cellular immunity efficiently. It is safe even after continuous administration and has a pleasant flavor. Dwg.0/0 FS CPI FA AR MC CPI: B14-A01; B14-A02; B14-A03; B14-A04; B14-A05; B14-G01; D03-H01T L108 ANSWER 16 OF 26 WPIX (C) 2003 THOMSON DERWENT ΑN 1999-163181 [14] WPIX DNC C1999-047550 TТ Skin activating agents and allergy inhibitory agents - comprise · fucoidan extracted from marine algae as effective component, exhibit e.g. hyaluronic acid biosynthesis inhibitory activities. DC B04 D21 (LIOY) LION CORP PΑ CYC 1 PT JP 11021247 A 19990126 (199914)\* 14p A61K035-80 ADT JP 11021247 A JP 1997-189249 19970630 PRAI JP 1997-189249 19970630 IC ICM A61K035-80 A61K007-00; A61K007-48; A61K031-725; C08B037-00 ICS JP 11021247 A UPAB: 19990412 AB Skin activating agents contain fucoidan, as effective component, extracted from one or more of marine algae belonging to Analipus, Nemacystis, Ecklonia, Lessonia, Macrocystis, Fucus, Ascophyllium, and Durvillea genera in Phaeophyceae. Also claimed are allergy inhibitory agents containing fucoidan, as effective component, extracted from one or more of marine algae belonging to Analipus, Nemacystis, Ecklonia, Lessonia, Macrocystis, Fucus, Ascophyllium, and Durvillea genera in Phaeophyceae. USE - The agents show potent hyaluronic acid biosynthesis and hyaluronidase inhibitory activities. The allergy inhibitory agents exhibit excellent histamine releasing inhibitory activity. Dwg.0/0 FS CPI FΑ AB MC CPI: B04-A08; B04-A10; B14-D07B; B14-G02A; B14-L09; B14-N17; D08-B09A L108 ANSWER 17 OF 26 WPIX (C) 2003 THOMSON DERWENT ΑN 1998-462862 [40] WPIX DNC C1998-140255 TΙ Production of highly pure fucoidan from cultured Nemacystus decipiens Kuck - by extraction with acid and treatment with e.g. diatomaceous earth. DC B04 D13 D21 PΑ (TAKO-I) TAKO M CYC 1 ΡI JP 10195106 A 19980728 (199840)\* 3р C08B037-00 JP 10195106 A JP 1997-36854 19970113 ADT PRAI JP 1997-36854 19970113 IC ICM C08B037-00 A61K031-725 ICA JP 10195106 A UPAB: 19981008 AB Production of fucoidan from cultured Nemacystus decipiens Kuck,

comprises extraction with HCl, H2SO4 or oxalic acid, esp. with lyophilisation, optionally with further purification of crude fucoidan by dissolution in BaCl2 solution and treatment with diatomaceous earth. ADVANTAGE - Low cost production of pure fucoidan useful for medicines, healthy foods and cosmetics. Dwg.0/0 CPI FS FΑ ΑB CPI: B04-C02; B04-F01; D03-H01T2; D08-B MC L108 ANSWER 18 OF 26 WPIX (C) 2003 THOMSON DERWENT 1998-460074 [40] WPIX AN DNC C1998-139074 Improvement in quality of Mozuku extract - comprises treating raw Mozuko TIor its extract with hydrogen peroxide. DC B04 D13 (HONS) YAKULT HONSHA KK PA CYC 1 PΙ A 19980728 (199840)\* 5p A23L001-337 JP 10191940 JP 10191940 A JP 1997-17932 19970116 ADT 19970116 PRAI JP 1997-17932 ICM A23L001-337 TC ICS A23L001-221; A23L001-30; A61K035-80 A61K031-715; C08B037-00 ICA JP 10191940 A UPAB: 19981008 AB Improvement in the quality of Mozuku extract containing fucoidan , comprises treating raw Mozuku, or its extract, with hydrogen peroxide. Also claimed are: (A) a process as above, followed by purification to remove low molecular weight impurities; and (B) an improved Mozuku extract produced by the above processes. ADVANTAGE - The process allows the colouration, characteristic odour and taste of Mozuku extract to be removed or reduced. Dwg.0/0 FS CPI FA AB; DCN MC CPI: B04-A07; B04-A10; B04-B02C; B05-C08; D03-H01; D03-H01L; D03-H01T2 L108 ANSWER 19 OF 26 WPIX (C) 2003 THOMSON DERWENT AN 1998-406047 [35] WPIX DNC C1998-122260 ΤI Foods containing fucoidan, separated from brown algae - e.g. confectionery, noodles, seasoning, pickles, cans, processed fruits, fish products and diary products, used to promote health. DC D13 PΑ (ITAY-I) ITAYA Y CYC 1 бp PΙ JP 10165114 A 19980623 (199835)\* A23L001-05 B2 19990809 (199937) **a**6 A23L001-05 JP 2932170 ADT JP 10165114 A JP 1996-357434 19961206; JP 2932170 B2 JP 1996-357434 19961206 FDTJP 2932170 B2 Previous Publ. JP 10165114 PRAI JP 1996-357434 19961206 IC ICM A23L001-05 A21D002-18; A23B007-10; A23G001-00; A23G003-00; A23G009-02; A23L001-20; ICA A23L001-22; A23L001-325; C08B037-00 10165114 A UPAB: 19980904 AΒ JΡ Foods containing fucoidan are new. The fucoidan is separated from brown algae and purified. USE - The foods are typically confectionery, noodles, seasoning, pickles, cans, processed fruits, fish products, diary products, health foods, processed drinks and liquor.

ADVANTAGE - The foods have good flavour, high safety and promote health. Dwg.0/0 CPI FS FΑ AB CPI: D03-H01T2 MC L108 ANSWER 20 OF 26 WPIX (C) 2003 THOMSON DERWENT 1998-234699 [21] WPIX AN C1998-073412 DNC TΤ Drugs for treatment of allergic diseases - comprising fucoidan or analogue prepared from sea weed. DC. (KYOD) KYODO NYUGYO KK PΑ CYC 1 JP 10072362 A 19980317 (199821)\* . 4p A61K035-80 PT JP 10072362 A JP 1996-245441 19960829 ADT 19960829 PRAI JP 1996-245441 ICM A61K035-80 IC A61K031-715 ICS AΒ JP 10072362 A UPAB: 19980528 Drugs for treatment of allergic diseases comprising fucoidan or an analogue prepared from sea weed are new. USE - The drugs can prevent the production of interleukin 4, immunoglobulin E, or histamine from mast cells. Dwg.0/0 FS CPI FΑ AB; DCN CPI: B04-A07F2; B04-A10; B04-D01; B05-C05; B07-A02; B14-G02A MC L108 ANSWER 21 OF 26 WPIX (C) 2003 THOMSON DERWENT 1998-172064 [16] ΑN WPIX DNC C1998-055104 TICancer therapeutic immune food - comprises U-fucoidan, D-fraction, beta-glucan, organic germanium and polysaccharide(s). DC B04 D13 PΑ (MATO-I) MATOBA J; (SOGA-I) SOGABE T CYC 1 A23L001-30 ΡI JP 10033142 A 19980210 (199816)\* 4p ADT JP 10033142 A JP 1996-225814 19960724 PRAI JP 1996-225814 19960724 IC ICM A23L001-30 A61K031-28; A61K031-555; A61K031-715; A61K035-80; A61K035-84 ICS AB JP 10033142 A UPAB: 19980421 Cancer therapeutic immune food is composed of combinations of several components with different pharmaceutical effects for synergism and acceleration of apoptosis and eradication of cancer cells containing Ufucoidan, D-fraction, beta -glucan, an organic germanium, and other polysaccharides. Dwg.0/3FS CPI FA AB; DCN CPI: B04-C02; B05-A02; B14-H01; B14-S09; **D03-H01T2** L108 ANSWER 22 OF 26 WPIX (C) 2003 THOMSON DERWENT 1998-051945 [05] WPIX DNC C1998-017767 TT Food and drinks containing fucoidan originating from fucoidan-containing substances such as seaweed - having apoptosis-inducing effects and being useful for preparing health foods, health drinks having anti-carcinogenic effects, stomach-controlling effects etc.. DC B03 D13 D16

```
IN
     IKAI, K; KATO, I; KIHARA, H; UMEDA, Y
     (TAKI) TAKARA SHUZO CO LTD; (IKAI-I) IKAI K; (KATO-I) KATO I;
PΑ
     (KIHA-I) KIHARA H; (UMED-I) UMEDA Y
CYC · 36
     WO 9747208
                   A1 19971218 (199805)* JA
                                              76p
PΙ
                                                     A23L001-30
        RW: AT BE CH DE DK EA ES FI FR GB GR IE IT LU MC NL PT SE
         W: AU BG BR CA CN CZ HU JP KR MX NO NZ PL RO SK US VN
                   A 19980107 (199820)
     AU 9727898
                                                     A23L001-30
     EP 916269
                   A1 19990519 (199924)
                                        EN
                                                     A23L001-30
         R: AT BE CH DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE
     CN 1221320
                   A 19990630 (199944)
                                                     A23L001-30
                     19990824 (199944)
     JP 10501425
                   Χ
                                                     A23L001-30
     AU 711896
                   В
                     19991021 (200002)
                                                     A23L001-30
     KR 2000010670 A 20000225 (200102)
                                                     A23L001-30
     US 2002076431 A1 20020620 (200244)
                                                     A61K035-80
    WO 9747208 A1 WO 1997-JP1664 19970515; AU 9727898 A AU 1997-27898
ADT
     19970515; EP 916269 A1 EP 1997-922085 19970515, WO 1997-JP1664 19970515;
     CN 1221320 A CN 1997-195371 19970515; JP 10501425 X WO 1997-JP1664
     19970515, JP 1998-501425 19970515; AU 711896 B AU 1997-27898 19970515; KR
     2000010670 A WO 1997-JP1664 19970515, KR 1998-708652 19981028; US
     2002076431 Al Cont of WO 1997-JP1664 19970515, Cont of US 1998-180465
     19981109, US 2001-987715 20011115
    AU 9727898 A Based on WO 9747208; EP 916269 A1 Based on WO 9747208; JP
FDT
     10501425 X Based on WO 9747208; AU 711896 B Previous Publ. AU 9727898,
     Based on WO 9747208; KR 2000010670 A Based on WO 9747208
PRAI JP 1996-318598
                      19961115; JP 1996-171666
                                                 19960612
IC
     ICM A23L001-30; A61K035-80
     ICS
         A61K031-715; A61K047-00; C07H005-10
ΑB
     WO
          9747208 A UPAB: 19980202
     Food or drinks contain apoptosis-inducing fucoidan originating
     from fucoidan-containing substances.
          USE - The food and drinks containing fucoidan are useful
     for apoptosis-induction and for health foods such as anti-carcinogenic
     foods and stomach/intestine caring foods.
          ADVANTAGE - The fucoidan originating from seaweed contains
     low or little alginic acid, the fucoidan does not disturb the
     original taste, flavour, texture and properties of food.
     fucoidan is cheap and safe.
     Dwg.0/5
FS
     CPI
FΑ
MC
     CPI: B07-A02B; B07-A03; B14-E10; B14-H01; D03-H01G; D03-H01T2;
          D05-C
L108 ANSWER 23 OF 26 WPIX (C) 2003 THOMSON DERWENT
     1993-045231 [05]
                        WPIX
DNN
                        DNC C1993-020417
TI
     Use of a blocking agent which inhibits leukocyte homing receptor mediated
     binding - for treating, diagnosing and monitoring e.g. multiple sclerosis.
DC
     B05 S03
     GEOFFROY, J; HUANG, K; ROSEN, S; SINGER, M; GEOFFREY, J
IN
     (REGC) UNIV CALIFORNIA
PΑ
CYC
     1.7
PΙ
     WO 9300919
                   A1 19930121 (199305) * EN
                                              29p
                                                     A61K037-00
        RW: AT BE CH DE DK ES FR GB GR IT LU MC NL SE
         W: JP
                                                     A61K031-70
     US 5227369
                   A 19930713 (199329)
                                                     A61K031-70
     EP 593658
                   A1 19940427 (199417)
                                         EN
         R: AT BE CH DE DK ES FR GB GR IT LI LU MC NL SE
     JP 07505859
                   W 19950629 (199534)
                                                     A61K045-00
                                                     A61K031-70
     EP 593658
                   B1 19991222 (200004)
                                         EN
         R: AT BE CH DE DK ES FR GB GR IT LI LU MC NL SE
                                                     A61K031-70
     DE 69230468
                   E 20000127 (200012)
```

```
ADT WO 9300919 A1 WO 1992-US5836 19920713; US 5227369 A US 1991-727280
     19910711; EP 593658 A1 EP 1992-915758 19920713, WO 1992-US5836 19920713;
     JP 07505859 W WO 1992-US5836 19920713, JP 1993-501815 19920713; EP 593658
     B1 EP 1992-915758 19920713, WO 1992-US5836 19920713; DE 69230468 E DE
     1992-630468 19920713, EP 1992-915758 19920713, WO 1992-US5836 19920713
FDT EP 593658 A1 Based on WO 9300919; JP 07505859 W Based on WO 9300919; EP
     593658 B1 Based on WO 9300919; DE 69230468 E Based on EP 593658, Based on
     WO 9300919
PRAI US 1991-727280
                      19910711
    21Jnl.Ref; EP 153875; EP 184040; US 4294818; US 4618601; US 4752563; US
     4818686; US 4839276; US 4935343; US 4948726; US 4994466; US 5036102; US
     5089479
IC
         A61K031-70; A61K045-00
     ICM
         A61K031-715; A61K037-02; A61K038-00; A61K039-395; G01N033-53
     ICS
          9300919 A UPAB: 19931119
AΒ
     Treating a demyelinating disease in a patient is claimed, comprising
     administering a compsn. comprising a carrier and a blocking agent which
     inhibits lymphocyte homing receptor (LHR)-mediated binding of leukocytes
     to myelin, the blocking agent being present in an amt. to inhibit
     LHR-mediated adhesion. The blocking agent may be e.g. mannose-6-phosphate,
     fructose-1-phosphate, a fragment of fucoidin or the
     phosphomannan monoester core from Hansenula hostii (PPME), Sqp50, Sqp90,
     an immunoglobulin or an isolated LHR.
          Also claimed is a method of blocking LHR-mediated adhesion of
     leukocytes to myelin in a patient, comprising administering a compsn.
     comprising a carrier and a blocking agent which inhibits LHR-mediated
     binding.
          USE - The blocking agents selectively bind either LHR or the
     recognition determinant on myelin. They can be used in the diagnosis and
     treatment of demyelinating diseases such as multiple sclerosis (MS), acute
     disseminated encephalomyelitis, acute necrotising haemorrhagic
     encephalomyelitis and HIV associated myelopathy.
     Dwq.0/0
FS
     CPI EPI
FA
     AB; DCN
     CPI: B04-B04A6; B04-B04C6; B04-C02; B05-B01P; B12-C10; B12-E01; B12-E02;
MC
         B12-G01
     EPI: S03-E14H4
ABEO US
          5227369 A UPAB: 19931116
     Treating the demyelmating effect comprises admin. of a compsn. comprising
     a protein blocking agent which inhibits LHR-mediated binding of leukocytes
     to myelin and inhibits adhesion. Blocking agent comprises an
     extracellullar region of an endothelial cell surface glycoprotein or an
     immunoglobulin.
          USE/ADVANTAGE - Used for treating and diagnosing demyelimating
     disease e.g. multiple sclerosis.
     Dwg.0/0
L108 ANSWER 24 OF 26 WPIX (C) 2003 THOMSON DERWENT
ΑN
     1992-249859 [30]
                       WPIX
     1988-021447 [03]; 1990-083347 [11]; 1990-099223 [13]; 1991-266907 [36];
CR
     1991-267279 [36]; 1992-307822 [37]; 1992-390086 [47]; 1993-377394 [47];
     1994-255250 [31]; 1994-332830 [41]; 1996-049424 [05]; 1996-057710 [06];
     1997-076903 [07]
DNC C1992-111476
TΙ
     Targetting of therapeutic agents using polysaccharide(s) - by forming
     complex of therapeutic agent with polysaccharide which interacts with cell
     receptor so complex is internalised into cells by RME.
DC
     B04 D16
     GROMAN, E V; JOSEPHSON, L; JUNG, C; LEWIS, J M
ΙN
PA
     (ADMA-N) ADVANCED MAGNETICS INC
CYC 18
```

PΙ

WO 9211037

A2 19920709 (199230)\* EN

19p

A61K047-48

```
RW: AT BE CH DE DK ES FR GB GR IT LU MC NL SE
        W: CA JP NO
    EP 563249
                  A1 19931006 (199340) EN
                                                     A61K047-48
        R: AT BE CH DE DK ES FR GB GR IT LI LU MC NL SE
    JP 06503347
                  W 19940414 (199420)
                                                     A61K047-48
    ES 2059299
                  T1 19941116 (199501)
    WO 9211037
                  A3 19920806 (199511)
    EP 563249
                  B1 19970423 (199721)
                                        ΕN
                                              11p
                                                     A61K047-48
        R: AT BE CH DE DK ES FR GB GR IT LI LU MC NL SE
    DE 69125848 E 19970528 (199727)
                                                     A61K047-48
    ES 2059299
                  T3 19971001 (199746)
                                                     A61K047-48
    CA 2097589
                  C 19980505 (199829)
                                                     A61K047-48
                  B2 20021216 (200302)
                                               q8
    JP 3357362
                                                     A61K047-48
ADT WO 9211037 A2 WO 1991-US9368 19911213; EP 563249 A1 WO 1991-US9368
    19911213, EP 1992-902979 19911213; JP 06503347 W WO 1991-US9368 19911213,
    JP 1992-503177 19911213; ES 2059299 T1 EP 1992-902979 19911213; EP 563249
    B1 WO 1991-US9368 19911213, EP 1992-902979 19911213; DE 69125848 E DE
    1991-625848 19911213, WO 1991-US9368 19911213, EP 1992-902979 19911213; ES
    2059299 T3 EP 1992-902979 19911213; CA 2097589 C CA 1991-2097589 19911213;
    JP 3357362 B2 WO 1991-US9368 19911213, JP 1992-503177 19911213
FDT EP 563249 A1 Based on WO 9211037; JP 06503347 W Based on WO 9211037; ES
    2059299 T1 Based on EP 563249; EP 563249 B1 Based on WO 9211037; DE
    69125848 E Based on EP 563249, Based on WO 9211037; ES 2059299 T3 Based on
    EP 563249; JP 3357362 B2 Previous Publ. JP 06503347, Based on WO 9211037
PRAI US 1990-630017
                     19901219
    No-SR. Pub; 8. Jnl. Ref; EP 281809; WO 9001295
IC
    ICM A61K047-48
    ICS
         A61K045-00; A61K047-36; A61P015-00; A61P031-12; A61P043-00
AB
         9211037 A UPAB: 20030111
    Targetting a therapeutic agent to a specific population of cells comprises
    forming a complex of the therapeutic agent with a polysaccaride capable of
    interacting with a cell receptor and allowing the complex to be
    internalised into the cells by receptor mediated endocytoxis.
         Before degradation or modification, the polysaccharide has mol. wt of
    over 1,000 D. First step includes modifying a first polysaccharide (pref.
    by degradation) constituting a second polysaccharide which can interact
    with a cell receptor, and forming a complex of the therapeutic agent with
    the second polysaccharide. Pref. the polysaccharide is arabinogalactan
     (aq.) mannan or fucoidan. Pref. combinations of polysaccharide
    and therapeutic agent are arabinogalactan and Fe, arabinogalactan and
    methotrexate, arabinogalactan and folic acid, arabinogalactan and ara
    A-phosphate, arabinogalactan and 6 alpha-methylprednisoloe and
    arabinogalactan and trifluorothymidine.
         Colloidal FeO coated with arabinogalactan was prepd as
    arabinogalactan is cleared by the asialoglycoprotein receptor of
    hepatocytes. The presence of injected Fe in the liver and not in the
     spleen indicates that targetting of Fe to hepatocytes has been achieved.
    USE/ADVANTAGE - Delivery of therapeutic agents eg Fe, folic acid,
    methotrexate, ara A-phosphate, 6 alpha-methyl prednisolone,
     trifluorothymidine, a hormone e.g. corticosteroid, a gene, enzyme or
     liposome. Antiviral agents may be targetted to hepatocytes to treat
    hepatitis, Fe to remedy anaemia etc., Conc. of therapeutic agents is
     increased in tissues where they have beneficial actions and decreased in
     tissues where they have unwanted or toxic effects.
     Dwg. 0/1
FS
    CPI
    AB; DCN
FA
MC
    CPI: B01-B02; B04-B03; B04-C02D; B05-A03A; B06-D09; B12-G02; B12-H01;
         D05-A01A1; D05-H
```

Targetting a therapeutic agent to a specific population of cells comprises forming a complex of the therapeutic agent with a polysaccharide capable of interacting with a cell receptor and allowing the complex to be

ABEQ EP

563249 A UPAB: 19931129

internalised into the cells by receptor mediated endocytosis.

Before degradation or modification, the polysaccharide has mol. wt of over 1,000 D. First step includes modifying a first polysaccharide (pref. by degradation) constituting a second polysaccharide which can interact with a cell receptor, and forming a complex of the therapeutic agent with the second polysaccharide. Pref. the polysaccharide is arabinogalactan (ag.) mannan or fucoidin. Pref. combinations of polysaccharide and therapeutic agent are arabinogalactan and Fe, arabinogalactan and methotrexate, arabinogalactan and folic acid, arabinogalactan and are A-phosphate, arabinogalactan and 6 alpha-methylprednisolone and arabinogalactan and trifluorothymidine.

USE/ADVANTAGE - Delivery of therapeutic agents eg Fe, folic acid, methotrexate, are A-phosphate, 6 alpha-methyl prednisolone, trifluorothymidine, a hormone e.g. corticosteroid, a gene, enzyme or liposome. Antiviral agents may be targetted to hepatocytes to treat hepatitis, Fe to remedy anaemia etc., Conc. of therapeutic agents is increased in tissues where they have beneficial actions and decreased in tissues where they have unwanted or toxic effects.

ABEQ EP 563249 B UPAB: 19970522

A complex for use in targeting a therapeutic agent to a specific population of cells, which complex comprises a carrier capable of binding to an RME receptor and selected from the group consisting of a polysaccharide and a modification thereof, and the therapeutic agent covalently bonded to the carrier such that the therapeutic agent may be targeted to the RME receptor on a cellular target and internalised therewith.

Dwg.0/0

L108 ANSWER 25 OF 26 WPIX (C) 2003 THOMSON DERWENT

AN 1988-220179 [31] WPIX

DNC C1988-098273

TI Anti metastatic and/or antiinflammatory compsns. - contg. sulphated polysaccharide.

DC B04 C03

IN PARISH, C R; SNOWDEN, J M; SNOWDEN, J

PA (PARI-I) PARISH C R; (AUSU) UNIV AUSTRALIAN NAT

CYC 17

PI WO 8805301 A 19880728 (198831) \* EN 53p RW: AT BE CH DE FR GB IT LU NL SE

W: AU JP US

AU 8812410 A 19880810 (198845) EP 355088 A 19900228 (199009) EN R: AT BE CH DE FR GB IT LI LU NL SE

JP 02502006 W 19900705 (199033) CA 1316828 C 19930427 (199322)

IL 85145 A 19940826 (199435) A61K031-725 IL 106354 A 19941021 (199443) A61K031-715 EP 631784 A1 19950104 (199506) EN 12p A61K031-725

A61K031-725

R: AT BE CH DE FR GB IT LI LU NL SE EP 355088 A4 19900816 (199512)

US 5541166 A 19960730 (199636) 12p A61K031-725 EP 355088 B1 19971210 (199803) EN 15p A61K031-725

R: AT BE CH DE FR GB IT LI LU NL SE JP 2701904 B2 19980121 (199808)

 JP 2701904
 B2 19980121 (199808)
 13p
 A61K031-725

 DE 3856083
 G 19980122 (199809)
 A61K031-725

 JP 09328431
 A 19971222 (199810)
 20p
 A61K031-725

 SG 45156
 A1 19980116 (199811)
 A61K031-725

 JP 2752353
 B2 19980518 (199825)
 19p A61K031-725

 SG 52246
 A1 1.9980928 (199903)
 A61K031-725

 EP 631784
 B1 19990331 (199917)
 EN A61K031-725

R: AT BE CH DE FR GB IT LI LU NL SE DE 3856321 G 19990506 (199924) A61K031-725

ADT WO 8805301 A WO 1988-AU17 19880122; EP 355088 A EP 1988-901519 19880122;

```
JP 02502006 W JP 1988-501413 19880122; CA 1316828 C CA 1988-557015
     19880121; IL 85145 A IL 1988-85145 19880120; IL 106354 A IL 1988-106354
     19880120; EP 631784 Al Related to EP 1988-901519 19880122, EP 1994-103441
     19880122; EP 355088 A4 EP 1988-901519
                                                  ; US 5541166 A Cont of WO
     1988-AU17 19880122, Cont of US 1989-391581 19890922, US 1992-853346
     19920316; EP 355088 B1 EP 1988-901519 19880122, WO 1988-AU17 19880122,
     Related to EP 1994-103441 19880122; JP 2701904 B2 JP 1988-501413 19880122,
     WO 1988-AU17 19880122; DE 3856083 G DE 1988-3856083 19880122, EP
     1988-901519 19880122, WO 1988-AU17 19880122; JP 09328431 A Div ex JP
     1988-501413 19880122, JP 1997-47706 19880122; SG 45156 A1 SG 1996-766
     19880122; JP 2752353 B2 Div ex JP 1988-501413 19880122, JP 1997-47706
     19880122; SG 52246 A1 SG 1996-1249 19880122; EP 631784 B1 Div ex EP
     1988-901519 19880122, EP 1994-103441 19880122; DE 3856321 G DE
     1988-3856321 19880122, EP 1994-103441 19880122
FDT
    IL 106354 A Div ex IL 85145; EP 355088 B1 Related to EP 631784, Based on
     WO 8805301; JP 2701904 B2 Previous Publ. JP 02502006, Based on WO 8805301;
     DE 3856083 G Based on EP 355088, Based on WO 8805301; JP 2752353 B2
     Previous Publ. JP 09328431; EP 631784 B1 Div ex EP 355088; DE 3856321 G
     Based on EP 631784
                      19870123; AU 1988-12410
                                                 19870113
PRAI AU 1987-9991
    1.Jnl.Ref; AU 8322582; AU 8430806; EP 140781; EP 165569; EP 208623; EP
     25123; GB 1029034; JP 60174729; US 4710493; 4.Jnl.Ref; EP 251134; EP
     254067; 9.Jnl.Ref; JP 61057520; WO 8807060
     A61K031-72; A61K045-05; C12N009-99
TC.
     ICM A61K031-715; A61K031-725
     ICS A61K031-72; A61K045-05; C12N009-99
ICA
    C08B037-10
ΑB
     WO
          8805301 A UPAB: 19930923
     Antimetastatic and/or antiinflammatory compsns. contain a sulphated
     polysaccharide (I) which blocks or inhibits endoglycosidase activity.

    is a heparinase-inhibiting sulphated polysaccharide, pref.

     fucoidan, pentosan sulphate, dextran sulphate, lambda-carrageenan
     or esp. heparin. The heparin may be modified to reduce its anticoagulant
     activity, esp. by decarboxylation or redn. and periodate oxidn.
          USE/ADVANTAGE - (I) inhibit metastasis of tumours (e.g. 13762 MAT
     mammary adenocarcinoma in rats), apparently by inhibiting passage of
     tumour cells through blood vessel walls, and are also active against
     experimental allergic encephalomyelitis.
     0/2
FS
     CPI
FΑ
     AB: DCN
     CPI: B04-C02; B12-A06; B12-C10; B12-D02; B12-D07; B12-G01B3;
MC
          B12-G07; C04-C02; C12-A06; C12-C10; C12-D02; C12-D07;
          C12-G01B3; C12-G07
          5541166 A UPAB: 19960913
ABEQ US
     A method of anti-metastatic treatment of an animal or human patient in
     need of such treatment, which comprises administration to the patient an
     anti-metastatic effective amount of sulphated polysaccharide which blocks
     or inhibits endoglycosidase activity, said sulphated polysaccharide being
     periodate-oxidized, reduced heparin.
     Dwg.0/2
           355088 B UPAB: 19980119
ABEQ EP
     Antimetastatic and/or antiinflammatory compsns. contain a sulphated
     polysaccharide (I) which blocks or inhibits endoglycosidase activity.
          (I) is a heparanase-inhibiting sulphated polysaccharide, pref.
     fucoidan, pentosan sulphate, dextran sulphate, lambda-carrageenan
     or esp. heparin. The heparin may be modified to reduce its anticoagulant
     activity, esp. by decarboxylation or redn. and oeriodate oxidn.
          USE/ADVANTAGE - (I) inhibit metastasis of tumours (e.g. 13762 MAT
     mammary adenocarcinoma in rats), apparently by inhibiting passage of
     tumour cells through blood vessel walls, and are also active against
     experimental allergic encephalomyelitis.
```

Dwg.0/2

```
L108 ANSWER 26 OF 26 WPIX (C) 2003 THOMSON DERWENT
     1985-130814 [22]
                        WPIX
ΑN
DNC C1985-056692
     Low salt content tangle prepn. - by immersing dry sea tangle in acetic
TΙ
     acid soln., maturing, washing, drying, re-immersing in acetic acid, adding
     seasoning etc..
DC
     D13
     (FUJI-N) FUJI KONBU KK
PA
CYC
ΡI
     JP 60066961
                   A 19850417 (198522)*
                                               5p
                   B 19860925 (198643)
     JP 61043030
     JP 60066961 A JP 1983-175586 19830922
ADT
PRAI JP 1983-175586
                      19830922
IC
     A23L001-33
         60066961 A. UPAB: 19930925
AΒ
     JΡ
     Method comprises (a) immersing starting dry sea tangle in aq. 8-15% acetic
     acid solns; (b) maturing; (c) washing in water at 40-100 deg.C for
     several-300 seconds favourably at 60-80 deg.C for 30-60 seconds; (d)
     drying to a moisture content of 5-30 (10-20)%; (e) immersing again in
     acetic acid soln; (f) maturing; (g) impregnating seasoning soln. contg.
     edible binder in it; and (h) preparing tangle flake as usual through
     pressing, maturing and shaving.
          Pref. the washing water contains at least one of alginic acid,
     (salt), aminoacids, laminaran, fucoidin, mannit, etc.; or the
     washing water is obtd. by de-salting once used washing water, to suppress
     the dissolution of taste and flavour components.
          USE/ADVANTAGE - Usually starting dry sea tangle has salt content
     6-12% and tangle flake prepd. from it, has salt content ca. 10%. Recently
     foods of low salt content have been required for health reasons. By
     washing sea tangle with water under specific conditions, salt can be
     dissolved out with suppressing the dissolution of taste and flavour
     components and tangle flake of salt content 4.5% can be prepd..
     0/0
FS
     CPI
FΑ
     AΒ
MC
     CPI: D03-H01T
=> d 1109 all abeg tech abex tot
L109 ANSWER 1 OF 14 WPIX (C) 2003 THOMSON DERWENT
AN
     2003-093322 [08]
                        WPIX
DNC
    C2003-023560
TI
     Drugs for preventing and/or treating fish infections due to viruses or
     bacteria, containing algae belonging to genera Kjellmaniella, Echlonia and
     Ascophyllum, or giant kelp or their extracts.
DC
     B04 C03
IN
     HAMASATO, K; KANEMITSU, A; KAWANO, T; NAGAOKA, M; NAKAO, M; OKUMURA, T;
     OMURA, H; SASAKI, M; UEYAMA, S; YAMASHITA, T; YOSHIMOTO, T
     (HAMA-I) HAMASATO K; (KANE-I) KANEMITSU A; (KAWA-I) KAWANO T; (MIYA-N)
PA
     MIYAKO KAGAKU CO LTD; (NAGA-I) NAGAOKA M; (NAKA-I) NAKAO M; (OKUM-I)
     OKUMURA T; (OMUR-I) OMURA H; (SASA-I) SASAKI M; (UEYA-I) UEYAMA S; (HONS)
     YAKULT HONSHA KK; (YAMA-I) YAMASHITA T; (YOSH-I) YOSHIMOTO T
CYC
     WO 2002092114 A1 20021121 (200308)* JA
ΡI
                                              23p
                                                     A61K035-80
         W: CN EC ID JP KR PH
     WO 2002092114 A1 WO 2002-JP4457 20020508
ADT
PRAI JP 2001-141272
                      20010511
IC
     ICM A61K035-80
     ICS A61K031-737; A61K035-74; A61P031-04; A61P031-12
AΒ
     WO 200292114 A UPAB: 20030204
     NOVELTY - Drugs (I) for preventing and/or treating fish infections contain
```

1 or more of algae belonging to the genera Kjellmaniella, Echlonia and Ascophyllum, giant kelp or their extracts.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) feeds for fish cultivation containing (I); and
- (2) a method for fish cultivation, or for preventing and/or treating fish infections, by administering (I) or the formulated feeds. ACTIVITY - Antibacterial; Virucide.

MECHANISM OF ACTION - None given in the source material.

USE - The drugs are for preventing and/or treating fish infections due to viruses or bacteria, particularly penaeid rod-shaped DNA virus (PRDV)-infection.

ADVANTAGE - The penaeid rod-shaped DNA virus (PRDV)-infection can be effectively prevented or treated.

Dwg.0/4

FS CPI

FA AB; DCN

MC CPI: B04-A08A; B04-A10; B04-F08; B04-F10B1; B14-A01; B14-A02; C04-A08A; C04-A10; C04-F08; C04-F10B1; C14-A01; C14-A02

TECH UPTX: 20030204

TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Drugs: (I) optionally contain **fucoidan**, and a bacterium of Bacillus subtilis or Bacillus lentus.

ABEX

ADMINISTRATION - (I) Are added to feeds during fish cultivation, at 60 - 80 mg daily.

EXAMPLE - Extracts of algae and giant kelp were obtained by boiling with water, centrifugation and drying the supernatant. Tests were carried out by using the extracts for formulating into feeds for fish cultivation, with high survival rate.

L109 ANSWER 2 OF 14 WPIX (C) 2003 THOMSON DERWENT

AN 2002-719667 [78] WPIX

DNC C2002-203818

TI Foodstuff, drink or fodder. e.g wheat-flour-, starch-, fats-and-oils-, soybean-processed products for human and animals e.g dog, fish, contains fucoidan, and agaro-oligosaccharide, as essential components.

DC D13

PA (TAKA-N) TAKARA BIO KK

CYC 1

PI JP 2002306131 A 20021022 (200278)\* 17p A23L001-308

ADT JP 2002306131 A JP 2001-117144 20010416

PRAI JP 2001-117144 20010416

IC ICM A23L001-308

ICS A21D002-18; A23C009-152; A23C011-10; A23C013-12; A23F003-14; A23G003-00; A23K001-16; A23L001-06; A23L001-16; A23L001-317; A23L001-325; A23L001-48; A23L002-52

AB JP2002306131 A UPAB: 20021204

NOVELTY - A foodstuff, drink or fodder contains fucoidan, its decomposition product, its salt, and an agaro-oligosaccharide.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for foodstuff, drink or fodder additive composition comprising fucoidan, its decomposition product, its salt, and an agaro-oligosaccharide.

ACTIVITY - Antioxidant; antiallergic; hemostatic; cytostatic; immunostimulant; antirheumatic; antiarthritic; antiinflammatory. 1 kg of **fucoidan** and 1 kg of agaro oligosaccharide was mixed with 136 kg of moisture fodder and administered to lymphocystis-diseased flat fishes at a dose of 0.1 g/kg body weight twice daily. The fodder efficiently treated lymphocystis disease and prevented relapse of lymphocystis disease.

MECHANISM OF ACTION - alpha -glucosidase inhibitor.

USE - Foodstuff (e.g. wheat-flour/starch processed products, noodles, bread, rice cake, fats-and-oils processed products, soybean processed products e.g tofu, fishery product e.g. boiled fish paste, sausage, dairy product e.g. cream and yogurt, vegetable-fruits processed product e.g. jam and vegetable/fruit drink, confectionery such as chocolates and biscuits, alcoholic beverages such as sake and wine; drink e.g. black tea, oolong eta and coffee, seasoning e.g. soy sauce and vinegar, etc) or fodder (e.g. stock raising product, processed marine product, etc) containing functional component derived from seaweed, for health maintenance. Also as fodder additive for improving health of pet animals e.g. dog, cat, horse or rabbit and marine animal e.g. oyster, shellfish, scallop, prawn, crab, etc.

ADVANTAGE - The foodstuff, drink and fodder has excellent physiological function improving effect and dietary fiber function, hence it provides excellent health improvement such as apoptosis induction, growth factor production potentiation, cytokine production regulation, antioxidant action, antiallergic effect, and hemostasis maintenance. Agaro-oligosaccharide prevents over production of nitrogen monoxide in the living body, provides antirheumatic-arthritic effect, antiinflammatory effect, inhibits endotoxin shock, inhibitor alpha -qlucosidase, etc. High functionality fodder, drink and foodstuff can be easily manufactured using the composition.

Dwg.0/0

FS CPI

FA AΒ

MC CPI: D01-A; D01-B; D03-C; D03-G; D06-A; D06-H; D06-H01

TECH UPTX: 20021204

TECHNOLOGY FOCUS - FOOD - Preferred Ingredients: The food, drink or fodder contains high concentration of components. The fucoidan is purified fucoidan. Agaro-oligosaccharide is agarobioses. Preferred Amount: The composition contains 0.1-99.9 (20-80) weight% (wt.%) each of fucoidan and agaro-oligosaccharide. Fucoidan and agaro-oligosaccharide are blended in the weight ratio of 0.1:99.9-99.9:0.1, preferably 30:70-70:30. Foodstuff and drinks contains 0.0001-40 (0.01-5) wt.% each of **fucoidan** and agaro-oligosaccharide.

ABEX

ADMINISTRATION - Fucoidan and oligosaccharide are each consumed at a dose of 0.0001-50 (0.01-10) mg/kg body weight daily, after adding the components into food or drink. Fucoidan and oligosaccharide are administered orally in powder form at a dose of 0.0001-2000 mg/kg body weight/day.

EXAMPLE - 7.3 kg of calcium chloride dihydrate was dissolved in 900 l of tap water. 20 g of dried product of Kjellmaniella tangle weed was ground and mixed with the water and temperature was raised from 12degreesC to 90degreesC by blowing water vapor for 40 minutes, subsequently heated at 90-95degreesC with stirring for 2 hours and cooled. 1100 l of cooled product was obtained. The cooled product was solid-liquid separated and 900 l of supernatant was obtained. 360 l of supernatant was concentrated to 20 1. Subsequently 20 1 of tap water was added and concentration step was repeated 5 times. Then desalting step was performed and 25 l of extract derived from Kjellmaniella tangle weed was obtained. 1 l of the solution was freeze dried and 13 g of Kjellmaniella tangle weed derived fucoidan dried product was obtained. Commercially available agar was dissolved in desalted water to produce 10% weight/volume solution. A strong cationic-exchange resin was added to the solution to produce a concentration of 1% weight/volume, then hydrolyzed at 90degreesC for 3 hours. The product was solid-liquid separated to remove resin from the solution. The filtrate was decolorized by adding 2 weight/volume% of activated carbon and filtered. The resulting solution was added to 1  $\mbox{N}$ sodium hydroxide and freeze-dried to produce agaro-oligosaccharide composition. The composition had pH of 5.2, moisture content of 2.3%, and contained galactose (9.8%), agarobiose (44.1%) and agaro-oligosaccharides except agarobiose (43.4%). A nutritive drink was prepared by compounding the tangle weed **fucoidan** and agaro-oligosaccharide. The nutritive drink was found to have excellent taste, fragrance and health improving effect.

```
L109 ANSWER 3 OF 14 WPIX (C) 2003 THOMSON DERWENT
     2002-404408 [43]
AN
                        WPIX
    C2002-113556
DNC
ΤI
     Use of fucoidan containing compositions for e.g. enhancing
     beta-transforming growth factor, preventing wrinkles, improving skin
     elasticity and promoting collagen production.
DC
     B04 D21
ΙN
     ADACHI, S; KATO, I; SAKAI, T; WU, H; YASUDA, M
PA
     (TAKI) TAKARA SHUZO CO LTD
CYC
     WO 2002006351 A1 20020124 (200243)* JA
PΤ
                                              66p
                                                     C08B037-00
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ
            NL OA PT SD SE SL SZ TR TZ UG ZW
         W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK
            DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KR KZ
            LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD
            SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW
     AU 2001069508 A 20020130 (200257)
                                                     C08B037-00
ADT WO 2002006351 A1 WO 2001-JP6032 20010712; AU 2001069508 A AU 2001-69508
     20010712
     AU 2001069508 A Based on WO 200206351
FDT
PRAI JP 2001-67445
                      20010309; JP 2000-212143
                                                 20000713; JP 2000-400615
     20001228
     ICM C08B037-00
IC
     ICS A61K007-00; A61K031-737; A61K035-56; A61K035-80; A61P017-00
AB
     WO 200206351 A UPAB: 20020709
     NOVELTY - Drugs or cosmetics comprising fucoidan, it's
     decomposition products or salts for:
          (1) treating or preventing diseases by enhancing beta -transforming
     growth factor ( beta -TGF);
          (2) ameliorating or preventing wrinkles;
          (3) increasing or maintaining skin elasticity;
          (4) ameliorating or preventing skin thickening;
          (5) preventing collagen reduction; and
          (6) enhancing collagen production.
          ACTIVITY - Dermatological. A hydrophilic ointment containing 25% v/v
     fucoidan extracted from Gagome using ethanol was applied to the
     backs of hairless mice at 250 ml/day five times a week and the mice were
     exposed to UVB radiation at thirteen weeks. After twenty-four weeks the
     skin thickness was 42 mm and skin contained 47.2 micro g/mg. The control
     results were 106 mm and 26 micro g/mg compared to 21 mm and 42 micro g/mg.
          MECHANISM OF ACTION - beta -TGF agonist; collagen agonist.
          USE - As a drug or cosmetic for:
          (1) treating or preventing diseases by enhancing beta -TGF;
          (2) improving or preventing wrinkles;
          (3) increasing or maintaining skin elasticity;
          (4) ameliorating or preventing skin thickening;
          (5) preventing collagen reduction; and
          (6) enhancing collagen production.
     Dwg.0/1
FS
     CPI
FΑ
     AB; DCN
     CPI: B04-C02D; B11-B; B14-L01; B14-N17; B14-R01; D08-B; D08-B09
MC
                    UPTX: 20020709
TECH
     TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Active Agent:
     Fucoidan is extracted from seaweed or Echinodermata species.
```

ADMINISTRATION - Dosage is 0.01-5 (preferably 0.1-2) g/day topically or

ABEX

0.1 mg-10 g (preferably 10 mg-2 mg) per day orally. L109 ANSWER 4 OF 14 WPIX (C) 2003 THOMSON DERWENT 2002-082793 [11] WPIX AN DNC C2002-025000 New fucoidan deacylase alpha-D-glucuronidase and TΙ endo-alpha-L-fucosidase for engineering sugar chains for anticancer and anticoagulant drugs. DC B04 D16 ΙN IKAI, K; ISHIZUKA, K; KATO, I; KOJIMA, K; SAKAI, T; SHIMANAKA, K PΑ (TAKI) TAKARA SHUZO CO LTD; (TAKI) TAKARA BIO INC CYC 95 WO 2001081560 A1 20011101 (200211)\* JA 145p C12N009-24 PΤ RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW C12N009-24 AU 2001048788 A 20011107 (200219) EP 1277834 A1 20030122 (200308) EN C12N009-24 R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI TR ADT WO 2001081560 A1 WO 2001-JP3333 20010419; AU 2001048788 A AU 2001-48788 20010419; EP 1277834 A1 EP 2001-921895 20010419, WO 2001-JP3333 20010419 AU 2001048788 A Based on WO 200181560; EP 1277834 Al Based on WO 200181560 PRAI JP 2000-186346 20000621; JP 2000-121116 20000421 IC ICM C12N009-24 ICS C08B037-00; C12N001-20; C12P019-04 ICA C12N015-11 ICI C12N015:11 WO 200181560 A UPAB: 20020215 AΒ NOVELTY - Fucoidan deacylase, alpha -D-glucuronidase and endoalpha -L-fucosidase of microbial origin, are new. DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following: (1) microorganisms producing the enzymes; (2) production of the enzymes by culture of the microorganisms; (3) preparation of deacetylated glucuronofucan sulfate, deacetylated glucuronofucan sulfate-containing oligosaccharides, and their decomposition products using the enzymes; (4) oligosaccharides prepared using the enzymes; and (5) activators for the enzymes. ACTIVITY - Cytostatic; anticoagulant. MECHANISM OF ACTION - Blood coaqulation reducer; apoptosis inducer. USE - The enzymes are used for engineering sugar chains to produce specific oligosaccharide structures active as anticoagulant and anticancer drugs. Dwg.0/40 FS CPI FA AB; DCN CPI: B04-C02X; B04-F01; B04-L01; B04-N03; B14-F04; B14-H01; D05-A04; MC D05-C03; D05-C08; D05-H08 TECH UPTX: 20020215 TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Enzyme: The fucoidan deacylase has a working pH of 6-9.1 and working temperature of 23-45 degreesC, and hydrolyses acetyl groups on glucuronofucan sulfate (or oligosaccharides containing it). The alpha-D-glucuronidase has a working pH 5.8-7.8 and working temperature 14-29 degreesC, and produces D-glucuronic acid by hydrolysis of the alpha-D-glucuronyl group on deacetylated glucuronofucan sulfate. The endo-alpha-L-fucosidase has a

working pH 4.5-7.5 and working temperature 23-42 degreesC, and hydrolyses

endo-alpha-L-fucosyl groups on deacetylated glucuronofucan sulfate.

**ABEX** 

AN

TΙ

DC

ΙN

PΑ

CYC PΙ

ICM C07K016-14; C07K016-44

ICS C07K017-00

DNC

Activators for the enzymes include sodium chloride, calcium chloride and proteins. The microorganism producing the enzymes is a Fucophilus strain having a genomic GC content of about 50 %. Glucuronofucan sulfate oligosaccharides include the specific structures of (I), (II), and (III). One or more of the free hydroxyl groups on the sugar residues bear sulfate or acetate groups, and all Fuc-Fuc linkages are (1-4). (Fuc-Fuc-Fuc-Fuc (2-GlucUA)) n-Fuc-Fuc-Fuc-Fuc Fuc-Fuc-Fuc-Fuc(2-GlucUA)-(Fuc-Fuc-Fuc-Fuc(2-GlucUA))n-Fuc-Fuc-Fuc-Fuc GlucUA(1-2)-Fuc-Fuc-Fuc-Fuc-Fuc (III) Fuc = L-fucose;GlucUA = D-glucuronic acid; and n = 1-5000. SPECIFIC MICROORGANISMS - Fucophilus fucoidanolyticus SI-1234 (FERM P-17517) is claimed. EXAMPLE - Fucophilus fucoidanolyticus SI-1234 (FERM P-17517) is cultured in the presence of 0.2 % glucuronofucan sulfate and 1 % peptone in artificial seawater medium. The cell bodies are disrupted by ultrasound and the supernatant purified on a diethylaminoethyl (DEAE)-Cellulofine (RTM) A-800 column with a 100mM to 40mM NaCl gradient of elution to isolate an enzyme fraction with 0.4 mU/ml activity. Use of this fraction to decompose glucuronofucan sulfate in 10 mM imidazole hydrochloride buffer containing 250 mM NaCl and 20 mM calcium chloride for 6 days at 30 degrees Centigrade and separation of the products on a DEAE-Cellulofine (RTM) A-800 column with a 20 mM to 600 mM NaCl elution gradient yields eight oligosaccharide products with molecular weights from 762 to 4216, and fucose/glucuronic acid molar ratios of 7:1 to 19:4. L109 ANSWER 5 OF 14 WPIX (C) 2003 THOMSON DERWENT 2002-082569 [11] WPIX C2002-024895 Antifucoidan antibody recognizing specific fucoidan structures, useful in functional studies and structural analysis of physiologically-active fucoidans of various origin as well as their quantizations for use in drugs and cosmetics. B04 D16 HINO, F; KATO, I; NAKAGAWA, K (TAKI) TAKARA SHUZO CO LTD WO 2000077049 A1 20001221 (200211)\* JA 27p C07K016-14 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TZ UG ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW AU 2000051061 A 20010102 (200216) C07K016-14 A1 20020313 (200225) ΕN C07K016-14 EP 1186616 R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI KR 2002007418 A 20020126 (200252) C07K016-14 CN 1354757 A 20020619 (200263) C07K016-14 JP 2001503905 X 20030114 (200316) C07K016-44 WO 2000077049 A1 WO 2000-JP3679 20000607; AU 2000051061 A AU 2000-51061 20000607; EP 1186616 A1 EP 2000-935565 20000607, WO 2000-JP3679 20000607; KR 2002007418 A KR 2001-714938 20011122; CN 1354757 A CN 2000-808570 20000607; JP 2001503905 X WO 2000-JP3679 20000607, JP 2001-503905 20000607 AU 2000051061 A Based on WO 200077049; EP 1186616 Al Based on WO 200077049; JP 2001503905 X Based on WO 200077049 PRAI JP 1999-165191 19990611

WO 200077049 A UPAB: 20020215 AΒ NOVELTY - Antifucoidan antibodies recognizing specific fucoidan structures, (I) and (II), are new. DETAILED DESCRIPTION - Antifucoidan antibodies recognizing specific fucoidan structures of formulae (I) and (II), are new. An INDEPENDENT CLAIM is also included for an antibody immobilized onto a support. USE - The antibodies are useful in the field of biochemistry in the functional studies and structural analysis of physiologically-active fucoidans of various origin, e.g. those with apoptosis induction activity, inhibitory activity on cancer proliferation and metastasis, antiviral activity and anticoagulation activity, as well as their quantizations for use in drugs and cosmetics. ADVANTAGE - Such antibodies can recognize fucoidan structures specifically. Dwq.0/1FS CPI AB; GI; DCN FA CPI: B04-C02; B04-C02X; B04-G01; B11-C07A; B12-K04A; D05-H09; D05-H11 MC TECH UPTX: 20020215 TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Antibody: Antibodies produced by GFD G-28 (FERM BP-7173) can preferably recognize (I) but not (II). Antibodies produced by GFD 2-9C (FERM BP-7174) can preferably recognize (II) but not (I). **ABEX** EXAMPLE - Fucoidans were isolated from kelp by the conventional method, which were injected into female Balb/c mice for immunization at 100 mg/ 100 ml then with an emulsion of incomplete fucoidan adjuvant. The spleen was homogenized for centrifugation, and the separated spleen cells were incubated with mouse myeloma cells P3U1 to give viable fusion cells for selection of antibody strains by immunoassay of the supernatant solutions. The two clones of GFD G-28 and GFD 2-9C producing cells were thus obtained for production of these antibodies. L109 ANSWER 6 OF 14 WPIX (C) 2003 THOMSON DERWENT 2000-217990 [19] WPIX AN DNC C2000-066672 TΤ Antibacterial agent, especially against Helicobacter pylori - for treating gastric ulcer contains fucoidan and/or its decomposed product. DC B04 D13 PA(TAKI) TAKARA SHUZO CO LTD CYC JP 2000044602 A 20000215 (200019)\* 5p PIC08B037-00 ADT JP 2000044602 A JP 1998-217282 19980731 PRAI JP 1998-217282 19980731 IC ICM C08B037-00 A23L001-30; A23L002-38; A23L002-52; A61K031-725 ICA A61K035-80 AΒ JP2000044602 A UPAB: 20000419 NOVELTY - An antibacterial agent contains fucoidan and/or its decomposed product as an active ingredient. USE - As antibacterial agent, particularly against Helicobacter pylori in foodstuffs or drinks (claimed). The antibacterial agent is used for maintaining health, as gastroenteric drink and to treat gastric ulcer. ACTIVITY - Antibacterial (especially against Helicobacter pylori (claimed). The antibacterial activity was measured by performing a growth inhibition test. The test was carried out by preparing various concentrations of fucoidan and/or its decomposed product (A,B and C) and were compared with purified water for the microbial count. The test samples and water were inoculated in 5% BHI broth medium containing fetal bovine serum at a pH of 7.3. The culture medium was incubated for

1-6 days in an anaerobic culture jar at 37 deg. C. The number of living

FS

FΑ

MC

ΤI

DC IN

PΑ

PI

IC

AB

FS :

FA

MC

microbes in test samples and water were measured in test samples and water were measured by surface smearing cultivation and calculated by counting CFU/ml (colony forming unit). The CFU were measured at 1,2,4 and 6 days and was found to be more in water (8.6). The concentration of test sample (A) was found to have dead microbes on fourth day itself which showed the effect of antibacterial effect especially against Helicobacter pylori. Dwg.0/1CPI AΒ CPI: B14-A01A; B14-E08; D03-H01G L109 ANSWER 7 OF 14 WPIX (C) 2003 THOMSON DERWENT 2000-013246 [01] WPIX DNC C2000-002517 New promoter for DNA synthesis reactions including polymerase chain reaction increases synthesis efficiency. ASADA, K; FUJITA, T; KATO, I; MIYAKE, K; MUKAI, H; SATO, Y; TAKEDA, O; UEMORI, T (TAKI) TAKARA SHUZO CO LTD CYC 78p WO 9954455 A1 19991028 (200001)\* JA C12N015-10 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SL SZ UG ZW W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KR KZ LC LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG US UZ VN YU ZA ZW A 19991108 (200014) AU 9935341 EP 1072678 A1 20010131 (200108) ENC12N015-10 R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE KR 2001042958 A 20010525 (200168) C12N015-10 CN 1306570 A 20010801 (200172) C12N015-10 JP 2000544787 X 20021105 (200304) C12N015-09 ADT WO 9954455 A1 WO 1999-JP2121 19990421; AU 9935341 A AU 1999-35341 ' 19990421; EP 1072678 A1 EP 1999-917081 19990421, WO 1999-JP2121 19990421; KR 2001042958 A KR 2000-711784 20001023; CN 1306570 A CN 1999-807709 19990421; JP 2000544787 X WO 1999-JP2121 19990421, JP 2000-544787 19990421 FDT AU 9935341 A Based on WO 9954455; EP 1072678 A1 Based on WO 9954455; JP 2000544787 X Based on WO 9954455 19981106; JP 1998-114005 PRAI JP 1998-315243 19980423 ICM C12N015-09; C12N015-10 ICS C120001-68 9954455 A UPAB: 20000105 NOVELTY - A promoter for DNA synthesis containing one or more acidic substances and/or cationic complexes is new. DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following: (1) methods of DNA synthesis using the promoter; and (2) kits for use in the DNA synthesis, including the promoter. USE - The promoter allows synthesis of DNA with higher efficiency than in conventional DNA synthesis reactions. Dwg.0/0 CPI AB; DCN CPI: B04-C02; B04-C03; B04-E02; B05-A03B; D05-H09; D05-H18B; D05-H19 TECH UPTX: 20000105 TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Promoter: This is an acidic substance such as an acidic polysaccharide or other high polymer, or a cationic complex such as a group VIII or other transition metal complex (especially of cobalt, iridium or rhodium). Preferred Method: DNA is amplified using the polymerase chain reaction, in

particular one using two or more DNA polymerases with 3'-5' exonuclease

activity (such as an alpha-DNA polymerase and a non-alpha non-pol I DNA polymerase).

**ABEX** 

SPECIFIC COMPOUNDS - The acidic substance used as promoter is fucoidan (including fucoidan-F and fucoidan -U), dextran sulphate, carragheenan, heparin, rhamnan sulphate, dermatan sulphate (chondroitin sulphate B), heparin sulphate, hyaluronic acid, alginic acid, pectin, polyglutamic acid, polyacrylic acid, polyvinyl sulphate, polystyrene sulphate or DNA. The cationic complex is (Co(NH3)6)Cl3, (Co(H2NCH2CH2NH2)3)Cl3 or (Rh(H2NCH2CH2NH2)3)Cl3.

EXAMPLE - 30 cycles of polymerase chain reaction are carried out using lambda-phage DNA (500pg) as template, with Pfu DNA polymerase I (1.25U), Pfu DNA polymerase II (1.25U) and three primer pairs designed to amplify fragments 1, 2 or 4 kb in length respectively from the template. The reaction is carried out in the presence of **fucoidan**-F (5ng or 50ng) as promoter, or for comparison in the absence of a promoter. Electrophoresis on the amplification product confirms amplification of the desired 1, 2 or 4 kb fragment when the promoter is present, and a higher amplification fragment yield for 50ng promoter than for 5ng promoter. When promoter is absent no detectable amplification occurs.

```
L109 ANSWER 8 OF 14 WPIX (C) 2003 THOMSON DERWENT
AN 1999-096182 [09] WPIX
DNC C1999-028503
```

TI New sugar compound comprising sulphation of at least one alcoholic hydroxy group - useful in medicine as anti-coagulant, anti-tumour and anti-AIDS virus infection etc..

DC B03 B04 D16 D17 E13

IN AKIYOSHI, S; IKAI, K; **KATO**, I; KIMURA, H; KOJIMA, K; NAKANISHI, Y; SAKAI, T

PA (REGL-N) RES INST GLYCOTECHNOLOGY; (TAKI) TAKARA SHUZO CO LTD CYC 1

PI AU 9724664 A 19981203 (199909)\* EN 135p C07H011-00

ADT AU 9724664 A AU 1997-24664 19970528

PRAI AU 1997-24664 19970528

IC ICM C07H011-00

ICS C08B037-00; C12N001-20; C12N009-24

AB AU 9724664 A UPAB: 19990302

A sugar compound represented by formula (I) or (II) are new comprising sulphation of at least one alcoholic hydroxyl group or its salt: X = H, (III); and Y = (IV), (V); Z = (VI); A1-6, B1-6 = H, SO3H. Also claimed are: (A) an endo-fucoidan-lyase having the following physicochemical properties: (i) acts upon fucoidan to liberate at least one sugar compound represented by the above; (ii) has optimum pH 6-10; and (iii) has optimum temperature 30-40 deg. C; and (B) a bacterium belonging to Fucoidanobacter which has menaquinone in the electron transport chain and contains approximately 60% of GC.

USE - The sulphated polysaccharide is useful in medicine as it has various biological activities e.g. anti-coagulant, lipaemia clearing, anti-tumour, cancer metastasis inhibitory and anti-AIDS virus infection effects. The sugar compounds are also useful in analysing the structure of fucoidan, identifying enzymatically degraded products of fucoidan and detecting the biological activities.

Dwg.0/41

FS CPI

FA AB; GI; DCN

MC CPI: B04-F1000E; B04-L06; B07-A02B; B10-A07; B14-A02B1; B14-F04; B14-F06; B14-H01; B14-H01B; D05-A02D; D05-H04; D05-H09; D06-G; E07-A02H; E07-A03C

L109 ANSWER 9 OF 14 WPIX (C) 2003 THOMSON DERWENT AN 1999-023989 [02] WPIX

```
DNC C1999-007249
DNN N1999-018484
     Agent for preventing infections in farmed fish and shellfish - contains
TΙ
     sulphated polysaccharide, especially fucoidan.
DC
     B04 C03 D13 P14
     KANEMITSU, A; NAGAOKA, M; OMURA, H; TAKAHASHI, Y; UEYAMA, S;
IN
     YAMASHITA, T; YOKOKURA, T
     (MIYA-N) MIYAKO KAGAKU KK; (HONS) YAKULT HONSHA KK
PA
CYC
     WO 9842204
                   A1 19981001 (199902)* JA
                                              15p
                                                     A23K001-16
PΤ
         W: CN ID JP KR
                  A 20000419 (200036)
     CN 1251020
                                                     A23K001-16
     JP 10545416
                   X 20000919 (200050)
                                                     A23K001-16
     KR 2000075860 A 20001226 (200134)
                                                     A23K001-16
ADT WO 9842204 A1 WO 1998-JP1145 19980318; CN 1251020 A CN 1998-803571
     19980318; JP 10545416 X JP 1998-545416 19980318, WO 1998-JP1145 19980318;
     KR 2000075860 A WO 1998-JP1145 19980318, KR 1999-707938 19990831
     JP 10545416 X Based on WO 9842204; KR 2000075860 A Based on WO 9842204
FDT
PRAI JP 1997-67973
                      19970321
     ICM A23K001-16
TC
     ICS A01K061-00; A23K001-18
AB
          9842204 A UPAB: 19990122
     Agent for prevention and treatment of infection in fish and shellfish
     contains a sulphated polysaccharide (I) as effective component.
          (I) is preferably fucoidan.
          USE - (I) is used in feedstuff for rearing fish and shellfish
     (claimed). (I) is used to protect e.g. prawns, yellow-tailed tuna,
     sea-bream and flat fish from infection with bacteria, viruses and
     parasites.
          ADVANTAGE - (I) protects from infection by iridoviridae.
     Dwq.1/2
FS
     CPI GMPI
FΑ
     AB; GI
     CPI: B04-C02; B14-E11; B14-S12; D03-G01
MC
L109 ANSWER 10 OF 14 WPIX (C) 2003 THOMSON DERWENT
     1999-018298 [02]
                       WPIX
AN
DNC
    C1999-005693
     Inhibitor of adhesion of Helicobacter pylori - contains effective
TΤ
     ingredient of glucuronic acid containing fucoidan, used for
     prevention of gastric and duodenal ulcers.
DC
PA
     (KAKE) KAKEN PHARM CO LTD; (TAKI) TAKARA SHUZO CO LTD
CYC
    1
PΙ
                 A 19981027 (199902)*
                                               q8
                                                     A61K031-725
     JP 10287571
     JP 10287571 A JP 1997-99237 19970416
ADT
                      19970416
PRAI JP 1997-99237
IC
     ICM A61K031-725
ICA A61K035-80
AR
     JP 10287571 A UPAB: 19990113
     An inhibitor of adhesion of Helicobacter pylori contains an effective
     ingredient of glucuronic acid containing fucoidan.
          USE - Used for prevention of gastric and duodenal ulcers.
          ADVANTAGE - Effective inhibition of gastritis, gastric and duodenal
     ulcers, including prevention of gastric cancer.
     Dwg.0/2
     CPI
FS
FΑ
     AB; DCN
MC
     CPI: B07-A02; B14-E08
L109 ANSWER 11 OF 14 WPIX (C) 2003 THOMSON DERWENT
AN
     1998-101040 [09]
                        WPIX
DNC C1998-033406
     Removal of cancer cells from compositions of haematopoietic cells - by
TΤ
```

treatment with apoptosis inducer such as fucoidan, dextran sulphate or 4,5-di hydroxy-2-cyclo penten-1-one. DC B04 D16 IN ASADA, K; KATO, I; KONISHI, H; KOYAMA, N PA (TAKI) TAKARA SHUZO CO LTD CYC 35 A1 19980115 (199809)\* JA WO 9801537 59p C12N005-00 PΙ RW: AT BE CH DE DK EA ES FI FR GB GR IE IT LU MC NL PT SE W: AU BG BR CA CN CZ HU JP KR MX NO NZ PL RO SK US VN AU 9732766 A 19980202 (199826) C12N005-00 A1 19990721 (199933) EP 930361 ΕN C12N005-00 R: DE FR GB IT NL A 19990728 (199948) CN 1224460 C12N005-00 X 19990921 (199950) C12N005-00 JP 10505035 KR 2000010744 A 20000225 (200102) C12N005-00 US 2001018209 A1 20010830 (200151) C12N005-08 WO 9801537 A1 WO 1997-JP2254 19970630; AU 9732766 A AU 1997-32766 ADT 19970630; EP 930361 A1 EP 1997-928515 19970630, WO 1997-JP2254 19970630; CN 1224460 A CN 1997-196137 19970630; JP 10505035 X WO 1997-JP2254 19970630, JP 1998-505035 19970630; KR 2000010744 A WO 1997-JP2254 19970630, KR 1998-708857 19981103; US 2001018209 A1 CIP of WO 1997-JP2254 19970630, CIP of US 1999-214609 19990108, US 2001-797821 20010305 AU 9732766 A Based on WO 9801537; EP 930361 A1 Based on WO 9801537; JP FDT 10505035 X Based on WO 9801537; KR 2000010744 A Based on WO 9801537 PRAI JP 1996-180500 19960710 IC ICM C12N005-00; C12N005-08 ICS C12N005-02 9801537 A UPAB: 19980302 AB WO Cancer cells are removed from compositions containing haematopoietic cells by treatment with an apoptosis inducing agent such as a sulphated polysaccharide or its degradation products (e.g. fucoidan or dextran sulphate), a carbohydrate containing uronic acid (or its derivatives or degradation products), or 4,5-dihydroxy-2-cyclopenten-1one. This treatment kills the cancer cells present in the composition without affecting the normal haematopoietic cells. USE - The method is used for production of haematopoietic cell compositions free from cancer cells, for use in treatment of cancer (such as myeloma) patients by in vitro treatment of marrow cells and for use in marrow transplantation. Compositions of haematopoietic cells into which foreign genes have been introduced (e.g. for gene therapy treatment of beta -thalassemia, sickling or recombinase deficiency) can also be freed from cancer cells by this method. Dwg.6/8 FS CPI FA AB; GI; DCN MC CPI: B04-C02; B04-F04; B10-E04A; B14-H01; D05-H08 L109 ANSWER 12 OF 14 WPIX (C) 2003 THOMSON DERWENT 1996-505810 [50] AN WPIX DNC C1999-028503 TΙ New sulphate ester(s) of tri, penta, or hexa saccharide cpds. - and new endo-fucoidan hydrolase and microorganism used for producing the sugars, useful as anticoagulants. DC B03 D16 AKIYOSHI, S; IKAI, K; KATO, I; KIMURA, H; KOJIMA, K; NAKANISHI, IN Y; SAKAI, T (REGL-N) RES INST GLYCOTECHNOLOGY; (TAKI) TAKARA SHUZO CO LTD; PΑ (AKIY-I) AKIYOSHI S; (IKAI-I) IKAI K; (KATO-I) KATO I; (KIMU-I) KIMURA H; (KOJI-I) KOJIMA K; (NAKA-I) NAKANISHI Y; (SAKA-I) SAKAI T CYC 25 A1 19961031 (199650) \* JA 113p PΙ WO 9634004 C07H011-00 RW: AT BE CH DE DK ES FI FR GB GR IE IT LU MC NL PT SE W: AT CA CN JP KR RU US

```
JP 08532343
                   X 19980630 (199836)
     EP 870771
                   A1 19981014 (199845)
                                         ΕN
         R: AT BE CH DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE
     AU 9724664
                   Α
                     19981203 (199909)
                                         EN 135p
                                                     C07H011-00
     KR 99008099
                   Α
                     19990125 (200014)
                                                     C07H011-00
     US 6054577
                   A 20000425 (200027)
                                                     C07H001-00
     CA 2217746
                   С
                      20000404 (200035)
                                         EN
                                                     C12N009-88
     US 6277616
                   B1 20010821 (200150)
                                                     C12N009-24
     AU 737081
                   B 20010809 (200152)
                                                     C07H011-00
     US 2001046696 A1 20011129 (200202)
                                                     C07H013-12
     CN 1330143
                   A 20020109 (200229)
                                                     C12N009-24
     US 6379947
                   B2 20020430 (200235)
                                                     C12N001-12
     CN 1183101
                   A 19980527 (200242)
                                                     C07H011-00
     RU 2193039
                   C2 20021120 (200307)
                                                     C07H011-00
ADT
    WO 9634004 A1 WO 1996-JP1080 19960422; JP 08532343 X JP 1996-532343
     19960422, WO 1996-JP1080 19960422; EP 870771 A1 EP 1996-910215 19960422,
     WO 1996-JP1080 19960422; AU 9724664 A AU 1997-24664 19970528; KR 99008099.
     A WO 1996-JP1080 19960422, KR 1997-707623 19971027; US 6054577 A WO
     1996-JP1080 19960422, US 1997-930002 19970926; CA 2217746 C CA
     1996-2217746 19960422, WO 1996-JP1080 19960422; US 6277616 B1 Div ex WO
     1996-JP1080 19960422, Div ex US 1997-930002 19970926, US 2000-517633
     20000303; AU 737081 B AU 1997-24664 19970528; US 2001046696 A1 Div ex WO
     1996-JP1080 19960422, Div ex US 1997-930002 19970926, Div ex US
     2000-517633 20000303, US 2001-893620 20010629; CN 1330143 A Div ex CN
     1996-193585 19960422, CN 2000-134824 19960422; US 6379947 B2 Div ex WO
     1996-JP1080 19960422, Div ex US 1997-930002 19970926, Div ex US
     2000-517633 20000303, US 2001-893620 20010629; CN 1183101 A CN 1996-193585
     19960422; RU 2193039 C2 WO 1996-JP1080 19960422, RU 1997-119864 19960422
     JP 08532343 X Based on WO 9634004; EP 870771 A1 Based on WO 9634004; KR
     99008099 A Based on WO 9634004; US 6054577 A Based on WO 9634004; CA
     2217746 C Based on WO 9634004; US 6277616 B1 Div ex US 6054577; AU 737081
     B Previous Publ. AU 9724664; US 2001046696 Al Div ex US 6054577, Div ex US
     6277616; US 6379947 B2 Div ex US 6054577, Div ex US 6277616; RU 2193039 C2
     Based on WO 9634004
PRAI JP 1995-127453
                      19950428
     JP 70059563; JP 70215990; JP 80000266
IC
     ICM
         C07H001-00; C07H011-00; C07H013-12; C12N001-12; C12N009-24;
          C12N009-88
     ICS
          C07H001-20; C07H005-10; C07H009-24; C07H013-02; C08B037-00;
          C12N001-20; C12P019-04; C12P019-14
ΑB
          9634004 A UPAB: 20010927
     Saccharide cpds. (A) and their salts are new. (A) are cpds. of formula (I)
     and (II) where at least one alcoholic hydroxyl qp. has been sulphated. A =
     a fucose gp. of formula (a); X = H \text{ or } A; Y = H \text{ or a qp. of formula (4)}
     bonded at a or b, with the other of a, b = OH; Z = H or a gp. of formula
     (6); provided that X and Y are not both H.
          Also claimed are (i) an endo-type fucoidan hydrolase (H)
     which releases a cpd. (7) and (8) from fucoidan, and has pH
     range 6-10 and temp. range 30-40 deg. C; and (ii) a Fucoidanobacter
     microorganism with menaquinone in the electron transmission chain and GC
     content of about 60%. (7) and (8) have formula (I; Y = H; X = A) with the
     3-OH of A sulphated in (7) and the 2 and 4 OH of A and the 5 OH of the
     galactose sulphated in (8). GC is not defined.
          USE - The sugar cpds and salts are useful for reagents for study of
     sugar chains. The new cpds. are useful as antitumour agents, metastasis
     inhibitors, and antiviral cpds. The endo-type fucoidan hydrolase
     and the Fucoidanobacter are useful for producing the new cpds.
          ADVANTAGE - The new cpds. are low mol. wt. cpds. and so, unlike
     fucoidan, are not antigenic.
     Dwg.0/42
FS
     CPI
```

CPI: B04-C02X; B04-F10; B04-L05B; B12-K04E; B14-A02B1; B14-F04; B14-H01;

FA

MC

AB; GI; DCN

D05-A02C; D05-C08; D05-H04; D06-G

```
L109 ANSWER 13 OF 14 WPIX (C) 2003 THOMSON DERWENT
     1996-379291 [38]
AN
                        WPIX
DNC C1996-119666
     Sulphuric ester of L-fucose useful as a contraceptive - is more effective
TΤ
     than use of spermicide e.g. poly oxyethylene nonyl phenyl-ether having no
     serious side effects by using oral contraceptive contg. estrogen and
     synthetic progesterone.
DC
     A96 B03 C02
     (TAKI) TAKARA SHUZO CO LTD; (TOSA-N) TOSA KOGAKU KENKYUSHO KK
PΑ
CYC
    1
PΙ
     JP 08183793 A 19960716 (199638)*
                                               4p
                                                     C07H011-00
     JP 08183793 A JP 1994-337597 19941228
ADT
PRAI JP 1994-337597
                      19941228
IC
     ICM C07H011-00
     ICS
        A61K031-70
     JP 08183793 A UPAB: 19960924
AB
     A saccharide deriv. comprises at least one OH qp. of L-fucose esterified
     with sulphuric acid.
          The L-fucose sulphuric ester is e.g. L-fucose-2-, 3-, 4- and
     5-sulphuric ester, L-fucose-2, 3- and 2, 4-disulphuric ester, L-fucose-2,
     3,4- and 2,3,5-trisulphuric ester, L-fucose-2,3,4,5 -tetrasulphuric ester,
     alpha-L-fucopyranosyl-2-sulphate-(1,2)-L- fucose and alpha-L-fucopyranosyl-
     2-sulphate-(1-2)-L-fucose. The sulphosaccharides can be obtd. pref. by
     acidic hydrolysis of natural fucoidan or fucane sulphate.
          USE/ADVANTAGE - A sulphuric ester of L-fucose is useful as a
     contraceptive. The sulphosaccharide inhibits a protein-polysaccharide
     interaction that closely relates to fertilisation. It acts more
     effectively than the use of a spermicide such as polyoxyethylene nonyl
     phenyl ether and does not have any serious side effect by the use of oral
     contraceptive contg. estrogen and synthetic progesterone.
          In an example, purified fucodaine is obtd. by purificn. of dried kelp
     extract (20 g.) was dissolved in 0.2 M aq. citric acid soln. and heated at
     pH 3 at 100 \ \text{deg.C} for 3 \ \text{hrs.} To the soln. was added 1 \ \text{M} aq. calcium
     acetate soln. (300 ml.) and the ppte. was removed. The filtrate was concd.
     and subjected to gel filtration chromatography to obtain a fraction of
     which the molecular wt. was up to 500. It was confirmed by high
     performance liq. chromatography that the fraction consisted of
     L-fucose-2-, 3-, 4- and 5-sulphuric ester and alpha-L-fucopranosyl-2-
     sulphate-(1-2)- L-fucose and alpha-L-fucopyranosyl-2-sulphate-(1-2)-L-
     fucose in a molar ratio of 4:5:5:10:6:4. The pretreatment of hamster
     spermatozoa with the sulphate L-fucose fraction or an hour completely
     inhibited fertilisation of matured hamster ova in vitro.
     Dwg.0/0
     CPI
FS
FΑ
     AB; DCN
     CPI: A12-W04C; B10-A07; C10-A07; B14-P01A; C14-P01A
MC
L109 ANSWER 14 OF 14 WPIX (C) 2003 THOMSON DERWENT
     1995-317479 [41]
AN
                        WPIX
DNC
    C1995-140937
ΤI
     Fucoidan oligosaccharide compsn. - used in cancer metastasis
     inhibiting agents and in antifungal agents..
DC
     (TAKI) TAKARA SHUZO CO LTD; (TOSA-N) TOSA KOGAKU KENKYUSHO KK
PA
CYC
PΙ
     JP 07215990
                  A 19950815 (199541)*
                                                q6
                                                     C07H003-06
ADT
     JP 07215990 A JP 1994-27589 19940201
PRAI JP 1994-27589
                      19940201
     ICM C07H003-06
     ICS
         A61K031-725
ICA A61K035-80
```

JP 07215990 A UPAB: 19951019 AΒ Fucoidan oligosaccharide compsn. has the following properties: (1) M.W. distribution: 5x103 or less (by a gel filtration through Cellulofine GCL-25); (2) Protein content: not detected; (3) Anti-aggregation activity: not held substantially. Also claimed are the prepn. of the fucoidan oligosaccharide compsn. in which fucoidan is hydrolysed by an organic acid under acid conditions; a cancer metastasis inhibiting agent contg. the  ${\tt fucoidan}$ oligosaccharide compsn.; and an antifungal agent contg. the fucoidan oligosaccharide compsn.. ADVANTAGE - The compsn. has good solubility and absorbability to living body, good reproducibility of biological activity and is also highly safe. In an example, 2g of fucoidan was dissolved in 100 ml water, the pH of the soln. was adjusted to 3.0 with acetic aid, and the soln. was held at 100 deg.C for 3 hrs.. The hydrolysate was gel filtered by Cellulofine CGL-25 column and the fraction of M.W. of 5,000 or lower was desalted and freeze-dried to give 1.98 g of prod. having no. anti-aggregation activity at 10 mg/ml or lower. Dwg.0/0 FS CPI FΑ AB CPI: B04-C02X; B14-A04; B14-H01B MC => fil frosti FILE 'FROSTI' ENTERED AT 10:11:20 ON 11 MAR 2003 COPYRIGHT (C) 2003 Leatherhead Food Research Association FILE LAST UPDATED: 10 MAR 2003 <20030310/UP> FILE COVERS 1972 TO DATE. => d all tot 1115 ANSWER 1 OF 11 FROSTI COPYRIGHT 2003 LFRA L115 FROSTI ΑN 590145 TΙ Remedies. ΤN Tominaga T.; Yamashita S.; Mizutani S.; Sagawa H.; Kato I. PΑ Takara Shuzo Co. Ltd SO European Patent Application PΙ EP 1226826 A1 20010301 WO 2001013925 20010301 20000817 ΑI PRAI Japan 19990820; 20000313 NTE 20010301 DTPatent LAEnglish э. SLEnglish AB Remedies for diseases needing the regulation of cytokines or allergic diseases contain fucoidan and/or its decomposition product as active ingredient. The formulations can induce the production of nitrogen monoxide. SH FUNCTIONAL FOODS CTALLERGIC DISEASES; ALLERGIES; CYTOKINES; EUROPEAN PATENT; FUCOIDAN; FUNCTIONAL FOODS; PATENT DED 23 Aug 2002 L115 ANSWER 2 OF 11 FROSTI COPYRIGHT 2003 LFRA ΑN 579386 FROSTI TΤ Homeostasis-maintaining agents. ΤN Nishiyama E.; Sagawa H.; Hino F.; Morihara E.; Sakai T.; Oyashiki H.; Kato I.

```
PA
      Takara Shuzo Co. Ltd
SO
      PCT Patent Application
PΙ
      WO 2002022140 A1
ΑI
      20010912
      Japan 20000913; 20000927; 20001109; 20001213; 20010425; 20010613
PRAI
DT
      Patent
      Japanese
LA
      English
SL
      Functional compositions and foods are described that contain
AΒ
      fucoidan or its decomposition products. They are said to help
      maintain homeostasis in humans and animals. Processes are disclosed for
      production of fucoidan and extracts of marine algae that have
      reduced colour, decreased bitterness, and lower iodine content.
      Applications include foods, beverages, seasonings, feeds, and drugs.
SH
      FUNCTIONAL FOODS
CT
      ALGAE; APPLICATIONS; EXTRACTION; FUCOIDAN; HOMEOSTASIS; MARINE
     ALGAE; PATENT; PCT PATENT
DED
      16 Apr 2002
     ANSWER 3 OF 11 FROSTI COPYRIGHT 2003 LFRA
L115
               FROSTI
ΑN
ΤI
      NUD (non-ulcer dyspepsia) - improving food.
IN
      Yoshikawa M.; Kudo T.; Nagaoka M.; Hashimoto H.; Kamiyama S.; Shibata H.;
      Takagi T.
PA
      Yakult Honsha Co. Ltd
SO
     Japanese Patent Application
PΙ
      JP 2001095528 A 20010410
AΙ
      19990927
NTE
      20010410
DT
     Patent
LA
     Japanese
SL
      English
AΒ
      A composition is described for a food that is claimed to be beneficial in
      the treatment of non-ulcer dyspepsia (NUD). The food contains extracts
      of a fucoidan derived from Phaeophyta, in addition to extracts
      such as senna tea, persimmon leaf, fennel or houttuynia cordata.
SH
      FUNCTIONAL FOODS
CT
      DISEASES; DYSPEPSIA; EXTRACTS; INTESTINAL DISORDERS; JAPANESE
     PATENT; NON ULCER DYSPEPSIA; PATENT; PLANT EXTRACTS;
      STOMACH ULCERS
DED
      15 Nov 2001
     ANSWER 4 OF 11 FROSTI COPYRIGHT 2003 LFRA
L115
ΑN
      564919
               FROSTI
TI
      Endo-fucoidan-lyase.
ΙN
      Sakai T.; Kimura H.; Kojima K.; Ikai K.; Akiyoshi S.; Nakanishi Y.; Kato
PA
      Takara Shuzo Co. Ltd; Research Institute for Glycotechology
SO
      United States Patent
PΙ
      US 6277616 B 20010821
ΑI
      20000303
NTE
      20010821
DT
      Patent
LA
      English
SL
      English
AΒ
      An endo-fucoidan-lyase is described that is useful in the
      production of sugar compounds for the study of carbohydrates. A
      microorganism of the genus Fucoidanobacter useful in the production of
      sugar compounds is also given. The lyase acts on fucoidan, and
      has a pH optimum of 6-10 and an optimum pH at 30-40 C.
SH
      PROCESSING
```

CARBOHYDRATES; ENDO FUCOIDAN LYASE; PATENT; SUGAR

COMPOUNDS; US PATENT

CT

DED 9 Oct 2001 L115 ANSWER 5 OF 11 FROSTI COPYRIGHT 2003 LFRA FROSTI ΤI Method and producing phosphorylated saccharides. IN Kamasaka H.; Okada S.; Kusaka K.; Yamamoto K.; Yoshikawa K. PΑ Ezaki Glico Co. Ltd SO United States Patent US 6268182 B 20010731 PΙ ΑI 19960925 PRAI Japan 19940811; 19950519 NTE 20010731 DT Patent LA English SLEnglish This patent continues from 5 861 048, which describes a method AB for producing phosphorylated saccharides. They can be used to stop alkali earth metals, e.g. calcium and iron, from precipitating, which helps their absorption by the body. These minerals have a positive effect on the health of the consumer, e.g. in preventing osteoporosis. The phosphorylated saccharides include a phosphate group that is obtained from glucan, mannan, dextran, agar, cyclodextrin, fucoidan, gellan gum, locust bean gum, guar gum, tamarind gum or xanthan gum. are said to prevent the formation of dental caries. FUNCTIONAL FOODS FUNCTIONAL INGREDIENTS; HEALTH FOODS; INGREDIENTS; MINERALS; CTPATENT; PHOSPHORYLATED SACCHARIDES; US PATENT DED 25 Sep 2001 L115 ANSWER 6 OF 11 FROSTI COPYRIGHT 2003 LFRA AN 551698 FROSTI TIRemedies. ΙN Inaga T.; Yamashita S.; Mizutani S.; Sagawa H.; Kato I. PΑ Takara Shuzo Co. Ltd SO PCT Patent Application PΤ WO 2001013925 A1 20010301 ΑI 20000817 PRAI Japan 19990820; 20000313 NTE 20010301 DTPatent LA English SLEnglish AΒ Remedies for diseases needing the regulation of cytokines or allergic diseases contain fucoidan and/or its decomposition product as active ingredient. The formulations can induce the production of nitrogen monoxide. SH FUNCTIONAL FOODS CTALLERGIC DISEASES; ALLERGIES; CYTOKINES; FUCOIDAN; FUNCTIONAL FOODS; PATENT; PCT PATENT DED 11 May 2001 ANSWER 7 OF 11 FROSTI COPYRIGHT 2003 LFRA L115AN 501824 FROSTI TT Foods or drinks. Umeda Y.; Kihara H.; Ikai K.; Kato I. IN PA Takara Shuzo Co. Ltd SO European Patent Application PΤ EP 916269 A1 ΑТ 19970515 PRAI Japan 19960612; 19961115 DΨ Patent

LA

English

SL AB Fucoidan is a polysaccharide containing sulfated fucose. occurs naturally in seaweed, and it has been shown to have apoptosis-inducing activity. This patent application covers foods and beverages containing fucoidan, and in which algins derived from the fucoidan-containing substance are reduced or removed. The patent outlines fucoidan extraction processes; active carbon can be used to remove the smell of seaweed. A very wide range of foods and beverages may be fortified with fucoidan. Among the examples quoted are processed cereal products, processed oil and fat products, soya products, meat products, dairy products, fruit and vegetable products, confectionery, bakery products, alcoholic beverages, infusion beverages, condiments, canned foods, bottled foods, convenience foods, dried foods, and frozen foods. An effective amount of fucoidan is added to a food or beverage to produce a product with apoptosis-inducing activity. SH FUNCTIONAL FOODS CTAPOPTOSIS; BEVERAGES; DIETARY SUPPLEMENTS; EUROPEAN PATENT; FORTIFIED BEVERAGES; FORTIFIED FOODS; FUCOIDAN; FUNCTIONAL BEVERAGES; FUNCTIONAL FOODS; FUNCTIONAL SUPPLEMENTS; NON ALCOHOLIC BEVERAGES; PATENT; SEAWEEDS DED 1 Sep 1999 L115 ANSWER 8 OF 11 FROSTI COPYRIGHT 2003 LFRA FROSTI ΤI Acetylfucoidan prepared from Okinawa Nemacystis decipiens and process for preparing the same. TN Tako M. SO PCT Patent Application PΙ WO 9901478 A1 ΑI 19980622 Japan 19970703; 19970909; 19980615 PRAI DT Patent LA Japanese SL English Fucoidans are polysaccharide products with interesting AΒ biological properties, such as anticancer activity, lipaemia-reducing activity, antitumour activity and anticoagulant activity. patent application relates to an acetylfucoidan prepared from algae or spores of Okinawa Nemacystis decipiens, and a method for its preparation. CTACETYLFUCOIDANS; ALGAL FUCOIDANS; ANTICANCER AGENTS; CHOLESTEROL LOWERING AGENTS; DIETARY SUPPLEMENTS; DIETETIC FOODS; FUCOIDANS; FUNCTIONAL FOODS; PATENT ; PCT PATENT DED 13 Apr 1999 L115 ANSWER 9 OF 11 FROSTI COPYRIGHT 2003 LFRA ΑN 488505 FROSTI TΙ Phosphorylated saccharide and method for producing the same. IN Kamasaka H.; Okada S.; Kusaka K.; Yamamoto K.; Yoshikawa K. PΑ Ezaki Glico Co. Ltd SO · United States Patent US 5861048 B 19990119 PΙ 19950811 ΑI Japan 19940811 PRAI 19990119 NTE DT Patent LA English

English This patent describes a method for producing phosphorylated AB saccharides. They can be used to stop alkali earth metals, e.g. calcium and iron, from precipitating, which helps their absorption by the body.

SL

These minerals have a positive effect on the health of the consumer, e.g. in preventing osteoporosis. The phosphorylated saccharides include a phosphate group that is obtained from glucan, mannan, dextran, agar, cyclodextrin, fucoidan, gellan gum, locust bean gum, guar gum, tamarind gum or xanthan gum. They are said to prevent the formation of dental caries.

CT FUNCTIONAL INGREDIENTS; HEALTH FOODS; INGREDIENTS; MINERALS; PATENT; PHOSPHORYLATED SACCHARIDES; US PATENT

DED 8 Mar 1999

L115 ANSWER 10 OF 11 FROSTI COPYRIGHT 2003 LFRA

AN 484221 FROSTI

TI Sugar compounds.

IN Sakai T.; Kimura H.; Kojima K.; Ikai K.; Akiyoshi S.; Nakanishi Y.; Kato

PA Takara Shuzo Co. Ltd

SO European Patent Application

PI EP 870771 A1

WO 9634004 19961031

AI 19960422

PRAI Japan 19950428

DT Patent

LA English

SL English

AB Fucoidan is a sulfated polysaccharide product of brown algae, and has a range of interesting physiological properties, including activity as an anticoagulant, lipaemia-reducing agent, anti-tumour and anti-cancer agent, and activity against the effects of AIDS virus infection. Fucoidan is very difficult to analyse and there is a need for the provision of known sugar compounds for use in analysing its structure and identifying its enzyme-degraded products, and characterization of their biological activities. This invention relates to sugar compounds and their salts represented by the illustrated general formula, in which at least one alcoholic hydroxyl group is sulfated. The second part of the invention relates to an endo-fucoidan-lyase, and the third part to a novel bacterium of the genus Fucanoidobacter useful for the production of such sugar compounds. Examples and details of production are given.

CT ANALYSIS; ENDOFUCOIDANLYASE; EUROPEAN PATENT; FUCANOIDOBACTER; FUCOIDAN; PATENT; POLYSACCHARIDES; SUGAR COMPOUNDS; SULFATED POLYSACCHARIDES

DED 12 Jan 1999

L115 ANSWER 11 OF 11 FROSTI COPYRIGHT 2003 LFRA

AN 460868 FROSTI

TI Foods or drinks.

IN Umeda Y.; Kihara H.; Ikai K.; Kato I.

PA Takara Shuzo Co. Ltd

SO PCT Patent Application

PI WO 9747208 A1

AI 19970515

PRAI Japan 19960612; 19961115

DT Patent

LA Japanese

SL English

AB The invention relates to functional or medical foods or drinks aimed at inducing apoptosis. The foods or drinks contain **fucoidan** obtained from **fucoidan**-containing substances, which have been treated to remove or reduce in quantity any algins present.

CT APOPTOSIS; FUCOIDAN; FUNCTIONAL FOODS;

MEDICAL FOODS; PCT PATENT

DED 10 Feb 1998

=> fil medline FILE 'MEDLINE' ENTERED AT 10:20:53 ON 11 MAR 2003

FILE LAST UPDATED: 8 MAR 2003 (20030308/UP). FILE COVERS 1958 TO DATE.

On June 9, 2002, MEDLINE was reloaded. See HELP RLOAD for details.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2003 vocabulary. See http://www.nlm.nih.gov/mesh/summ2003.html for a description on changes.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d all tot

L134 ANSWER 1 OF 3 MEDLINE

AN 2000300937 MEDLINE

DN 20300937 PubMed ID: 10841555

- TI Mobilization of stem/progenitor cells by sulfated polysaccharides does not require selectin presence.
- AU Sweeney E A; Priestley G V; Nakamoto B; Collins R G; Beaudet A L; Papayannopoulou T
- CS Department of Medicine, Division of Hematology, University of Washington, Seattle, WA 98195-7710, USA.
- NC AI32177 (NIAID) HL46557 (NHLBI) HL58734 (NHLBI)
- SO PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE UNITED STATES OF AMERICA, (2000 Jun 6) 97 (12) 6544-9.

  Journal code: 7505876. ISSN: 0027-8424.
- CY United States
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Priority Journals
- EM 200007
- ED Entered STN: 20000720 Last Updated on STN: 20000720 Entered Medline: 20000713
- Employing carbohydrate ligands, which have been extensively used to block AB selectin function in vitro and in vivo, we have examined the involvement of such ligands in stem/progenitor cell mobilization in mice and monkeys. We found that sulfated fucans, branched and linear, are capable of increasing mature white cells in the periphery and mobilizing stem/progenitor cells of all classes (up to 32-fold) within a few hours posttreatment in a dose-dependent manner. To elicit the effect, the presence of sulfate groups was necessary, yet not sufficient, as certain sulfated hexosamines tested (chondroitin sulfates A or B) were ineffective. Significant mobilization of stem/progenitor cells and leukocytosis was elicited in selectin-deficient mice (L(-/-), PE(-/-), or LPE(-/-)) similar to that of wild-type controls, suggesting that the mode of action of sulfated fucans is not through blockade of known selectins. Other mechanisms have been entertained, in particular, the release of chemokines/cytokines, including some previously implicated in mobilization. Significant increases were documented in the levels of seven circulating chemokines/cytokines within a few hours after fucan sulfate treatment and support such a proposition. Additionally, an increase was noted in plasma metalloproteinase (MMP) 9, which might independently contribute to the mobilization process by enzymatically facilitating chemokine/cytokine release. Mobilization by sulfated polysaccharides provides a distinct paradigm in the mobilization process and uncovers an

additional novel in vivo biological role for sulfated glycans. As similarly sulfated compounds were ineffective in vivo, the data also underscore the fact that polysaccharides with similar structures may elicit diverse in vivo effects.

CT Check Tags: Animal; Support, U.S. Gov't, P.H.S.

Chemokines: BL, blood Cytokines: BL, blood

Gelatinase B: ME, metabolism

\*Hematopoietic Stem Cell Mobilization

Macaca nemestrina

Mice

\*Polysaccharides: PD, pharmacology

\*Selectins: PH, physiology

Structure-Activity Relationship

RN 9072-19-9 (fucoidan)

CN 0 (Chemokines); 0 (Cytokines); 0 (Polysaccharides); 0 (Selectins); EC 3.4.24.35 (Gelatinase B)

L134 ANSWER 2 OF 3 MEDLINE

AN **97279844** MEDLINE

DN 97279844 PubMed ID: 9134218

TI The effect of the selectin binding polysaccharide **fucoidin** on eosinophil recruitment in vivo.

AU Teixeira M M; Hellewell P G

CS Imperial College School of Medicine, National Heart and Lung Institute, London.

SO BRITISH JOURNAL OF PHARMACOLOGY, (1997 Mar) 120 (6) 1059-66. Journal code: 7502536. ISSN: 0007-1188.

CY ENGLAND: United Kingdom

DT Journal; Article; (JOURNAL ARTICLE)

LA English

FS Priority Journals

EM 199708

ED Entered STN: 19970908 Last Updated on STN: 19970908 Entered Medline: 19970827

1. In order to accumulate at sites of inflammation, leukocytes initially AB roll on endothelial cells of postcapillary venules before becoming firmly attached. This process of rolling is mediated by selectins which bind to carbohydrate counter-ligands present on the surface of both leukocytes and endothelial cells. The polysaccharide fucoidin has been previously shown to inhibit leukocyte rolling in the mesenteric circulation and to reduce neutrophil accumulation in the skin and meninges in experimental inflammation. 2. In the present study we have assessed the effects of fucoidin on eosinophil function in vitro and eosinophil accumulation at sites of inflammation in guinea-pig skin. 3. At concentrations of up to 1200 micrograms ml-1, fucoidin inhibited phorbol myristate acetate (PMA)-induced eosinophil homotypic aggregation by up to 60% but had no inhibitory effect on PMA-induced eosinophil adhesion to serum-coated plates. 4. Fucoidin effectively reduced the binding of the anti-L-selectin mAb MEL-14 to guinea-pig eosinophils. Binding of a P-selectin-IgG chimera to eosinophils was also partially inhibited by fucoidin, but binding of an anti-CD18 or an anti-VLA-4 mAb were unaffected. 5. When given systemically to quinea-pigs, fucoidin suppressed 111In-labelled eosinophil recruitment to sites of allergic inflammation. 111In-labelled eosinophil accumulation induced by platelet-activating factor (PAF) and zymosan-activated plasma (as a source of C5a des Arg) was also inhibited. 6. These results demonstrate a role for fucoidin-sensitive selectins in mediating eosinophil recruitment in vivo.

CT Check Tags: Animal; Comparative Study; Support, Non-U.S. Gov't Antibodies, Monoclonal: IM, immunology Antibodies, Monoclonal: PD, pharmacology

```
*Antigens, CD18: IM, immunology
     *Chemotaxis, Leukocyte: DE, drug effects
      Chemotaxis, Leukocyte: IM, immunology
        Drug Eruptions: IM, immunology
     *Eosinophilia: IM, immunology
     *Eosinophils: DE, drug effects
      Eosinophils: IM, immunology
      Guinea Pigs
      Integrins: IM, immunology
     *Polysaccharides: PD, pharmacology
      Receptors, Lymphocyte Homing: IM, immunology
     *Selectins: IM, immunology
      Skin: DE, drug effects
     *Skin: IM, immunology
      Tetradecanoylphorbol Acetate
     16561-29-8 (Tetradecanoylphorbol Acetate); 9072-19-9 (fucoidan)
RN
CN
     0 (Antibodies, Monoclonal); 0 (Antigens, CD18); 0 (Integrins); 0
     (Polysaccharides); 0 (Receptors, Lymphocyte Homing); 0 (Selectins); 0
     (integrin alpha4beta1)
L134 ANSWER 3 OF 3
                       MEDLINE
ΑN
     97228143
                  MEDLINE
DN.
     97228143
               PubMed ID: 9091581
TI
     The association between alpha4-integrin, P-selectin, and E-selectin in an
     allergic model of inflammation.
ΑU
     Kanwar S; Bullard D C; Hickey M J; Smith C W; Beaudet A L; Wolitzky B A;
CS
     Department of Medical Physiology, University of Calgary, Alberta, Canada.
NC
     AI-32177 (NIAID)
     GM-15483 (NIGMS)
     HL-42550 (NHLBI)
SO
     JOURNAL OF EXPERIMENTAL MEDICINE, (1997 Mar 17) 185 (6) 1077-87.
     Journal code: 2985109R. ISSN: 0022-1007.
CY
     United States
DT
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
FS
     Priority Journals
EM
     199704
ED
     Entered STN: 19970422
     Last Updated on STN: 19970422
     Entered Medline: 19970410
     In this study, we examined the relationship between the endothelial
AΒ
     selectins (P-selectin and E-selectin) and whether they are critical for
     alpha4-integrin-dependent leukocyte recruitment in inflamed (late phase
     response), cremasteric postcapillary venules. Animals were systemically
     sensitized and 2 wk later challenged intrascrotally with chicken
     ovalbumin. Leukocyte rolling flux, adhesion, and emigration were assessed
     at baseline and 4 and 8 h postantiqen challenge. There was a significant
     increase in leukocyte rolling flux, adhesion, and emigration in sensitized
     and challenged mice at both 4 and 8 h. At 8 h, the increase in leukocyte
     rolling flux was approximately 50% inhibitable by an anti-alpha4-integrin
     antibody, 98% inhibitable by fucoidin (a selectin-binding
     carbohydrate), and 100% inhibitable by an anti-P-selectin antibody.
     P-selectin-deficient animals displayed no leukocyte rolling or adhesion at
     8 h after challenge. However, at 8 h there were many emigrated leukocytes
     in the perivascular space suggesting P-selectin-independent rolling at an
     earlier time point. Indeed, at 4 h postantigen challenge in
     P-selectin-deficient mice, there was increased leukocyte rolling,
     adhesion, and emigration. The rolling in the P-selectin-deficient mice at
     4 h was largely alpha4-integrin dependent. However, there was an essential
```

E-selectin-dependent component inasmuch as an anti-E-selectin antibody completely reversed the rolling, and in E-selectin and P-selectin double deficient mice rolling, adhesion and emigration were completely absent.

```
These results illustrate that P-selectin underlies all of the
     antigen-induced rolling with a brief transient contribution from
     E-selectin in the P-selectin-deficient animals. Finally, the
     antigen-induced alpha4-integrin-mediated leukocyte recruitment is entirely
     dependent upon endothelial selectins.
CT
     Check Tags: Animal; Support, Non-U.S. Gov't; Support, U.S. Gov't, P.H.S.
     *Antigens, CD: PH, physiology
      Cell Adhesion
      Chickens
      Crosses, Genetic
       *Hypersensitivity, Immediate: IM, immunology
        Hypersensitivity, Immediate: PP, physiopathology
      Immunization
     *Inflammation: IM, immunology
      Inflammation: PP, physiopathology
      L-Selectin: GE, genetics
     *L-Selectin: PH, physiology
      Leukocytes: IM, immunology
      Leukocytes: PH, physiology
      Mice
      Mice, Inbred C57BL
      Mice, Inbred Strains
      Mice, Knockout
      Ovalbumin: IM, immunology
      P-Selectin: GE, genetics
     *P-Selectin: PH, physiology
      Time Factors
RN
     126880-86-2 (L-Selectin); 143198-26-9 (alpha4 integrin); 9006-59-1
     (Ovalbumin)
     0 (Antigens, CD); 0 (P-Selectin)
CN
=> d his
     (FILE 'HOME' ENTERED AT 08:53:17 ON 11 MAR 2003)
                SET COST OFF
     FILE 'HCAPLUS' ENTERED AT 08:53:28 ON 11 MAR 2003
                E FUCOIDAN/CT
                E E3+ALL
            684 S E4
L1
     FILE 'REGISTRY' ENTERED AT 08:53:58 ON 11 MAR 2003
L2
              1 S 9072-19-9
L3
              5 S 9072-19-9/CRN
     FILE 'HCAPLUS' ENTERED AT 08:54:41 ON 11 MAR 2003
L4
           1028 S FUCOIDIN# OR FUCOIDAN# OR NEMACYSTUS MUCILAGE
L5
              5 S L3
L6
           1048 S L1, L4, L5
                E FUCOIDAN
1.7
            739 S E3
\Gamma8
            316 S E15, E17
L9
           1048 S L6-L8
L10
            225 S NITROGEN MONOOXIDE
     FILE 'REGISTRY' ENTERED AT 08:56:36 ON 11 MAR 2003
L11
              1 S 10102-43-9
     FILE 'HCAPLUS' ENTERED AT 08:57:40 ON 11 MAR 2003
          70393'S L11
L12
L13
         135853 S OHM11771 OR OHM()(11771 OR 11 771) OR NITROGEN()(MONOXIDE OR
L14
              7 S L10, L12, L13 AND L9
```

```
E INTERLEUKIN/CT
L15
           5859 S E39
                 E E144+ALL
           6645 S E23, E46
L16
          85930 S E7, E6+NT
L17
L18
         117789 S IL OR IL12 OR INTERLEUKIN OR (IL OR INTERLEUKIN) (L)12
L19
              38 S L9 AND L15-L18
                 E INTERFERON/CT
             300 S E3
L20
L21
          30764 S E89
                 E E71+ALL
          54117 S E6+NT
L22
          30764 S E6(L)GAMMA
L23
          41617 S IFNGAMMA OR GAMMAIFN OR (IFN OR INTERFERON) (L) GAMMA
L24
L25
              15 S L9 AND L20-L24
                 E IGE/CT
                 E E3+ALL
           9782 S E2
L26
                 E IMMUNOGLOBULIN/CT
                 E IMMUNOGLOBULINS/CT
L27
           9782 S E38, E39
                 E E3+ALL
          10193 S E7, E6 (L) "E"
L28
L29
               3 S L9 AND L26-L28
L30
               6 S L9 AND (IGE OR (IG OR IMMUNOGLOBULIN) (S) "E")
                 E CYTOKINE/CT
                 E E48+ALL
          76929 S E5,E4
L31
         157526 S E4+NT
L32
L33
             46 S L9 AND L31, L32
             37 S L9 AND CYTOKINE
L34
L35
              18 S L9 AND LYMPHOKINE
L36
              66 S L14, L19, L25, L29, L30, L33-L35
                 E WO2000-JP5489/AP, PRN
L37
               1 S E3, E4
                 E JP99-234262/AP, PRN
L38.
               1 S E4
                 E JP2000-69223/AP, PRN
               1 S E4
L39
                 E TAKARA/PA,CS
            772 S E93-E129
L40
           1582 S E3-E145
L41
           3405 S (TAKARA? OR SHUZO?)/PA,CS
L42
L43
             29 S L9 AND L40-L42
                 E TOMINAGA T/AU
L44
             218 S E3, E4, E17-E19
                 E TAKANARI/AU
                 E YAMASHITA S/AU
             385 S E3
L45
                 E YAMASHITA SYU/AU
               7 S E6, E7
L46
                 E SYUSAK/AU
                 E MIZUTANI S/AU
L47
             106 S E3, E34
                 E SHIGETOSHI/AU
                 E SAGAWA H/AU
             386 S E3, E11, E12
L48
                 E HIROAKI S/AU
L49
               1 S E3
                 E KATO I/AU
             728 S E3-E5, E22-E25
L50
                 E IKUNOSH/AU
L51
               5 S E4
```

```
L52
             35 S L44-L51 AND L9
L53
             2 S L36 AND L43, L52
             34 S L43, L52 NOT L53
L54
L55
             14 S L54 AND (FOOD# OR FEED? OR BEVERAGE# OR HEALTH FOOD# OR DRUG#
                SEL DN AN 3 6 10 13 14
L56
              9 S L55 NOT E1-E15
             0 S L54 AND (FOOD? OR NUTRI?)/SC, SX NOT L55
L57
L58
             64 S L36 NOT L40-L57
                SEL DN AN 12 14 16 27 30 40 52
              7 S L58 AND E16-E36
L59
                E ALLERGY/CT
                E E3+ALL
          19556 S E3, E2+NT
L60
                E E15+ALL
L61
           7639 S E3
                E E7+ALL
L62
           6554 S E4
                E E15+ALL
          13841 S E5
L63
                E E4+ALL
          31196 S E4+NT
L64
                E E13+ALL
           8958 S E4, E3+NT
L65
                E IMMUN/CT
                E IMMUNOS/CT
L66
          15342 S E12+NT OR E20+NT
          25407 S E26+NT OR E27+NT
L67
L68
             26 S L9 AND L60-L67
L69
             11 S L68 NOT L36, L40-L59
L70
             91 S L9 AND (NUTRI? OR FOOD? OR FEED? OR BEVERAG? OR DRINK? OR JUI
            79 S L9 AND (BEVERAG? OR ?DRINK? OR ?JUICE? OR FOOD? OR FEED?)/BI
L71
L72
            30 S L9 (L) FFD/RL
L73
            18 S L53, L56, L59
L74
             4 S L68 AND L73
L75
            18 S L73, L74
L76
            14 S L70-L72 AND L75
          . 18 S L75, L76
L77
L78
            89 S L70-L72 NOT L77
L79
             67 S L78 AND (PY<=2000 OR PRY<=2000 OR AY<=2000)
L80
             37 S L79 AND (FOOD? OR NUTRI?)/SC
                SEL DN AN 5 24
L81
             2 S L80 AND E1-E6
             20 S L77, L81
L82
             30 S L79 NOT L80
L83
L84
             74 S L9 AND (?INFLAM? OR LEUKOTRIEN?)
L85
             19 S L84 AND L19, L29, L30, L36, L60-L69
L86
             1 S L84 AND L70-L72
L87
             20 S L85, L86
                SEL DN AN 10 11 14 15 16 19 20
L88
              7 S E7-E27 AND L87
L89
             27 S L82, L88 AND L4-L10, L12-L88
     FILE 'REGISTRY' ENTERED AT 09:53:35 ON 11 MAR 2003
     FILE 'HCAPLUS' ENTERED AT 09:53:56 ON 11 MAR 2003
     FILE 'REGISTRY' ENTERED AT 09:55:26 ON 11 MAR 2003
L90
           1 S 328081-45-4
     FILE 'HCAPLUS' ENTERED AT 09:55:36 ON 11 MAR 2003
L91
              1 S L90
```

FILE 'USPATFULL, USPAT2' ENTERED AT 09:55:40 ON 11 MAR 2003

FILE 'REGISTRY' ENTERED AT 09:55:49 ON 11 MAR 2003

FILE 'HCAPLUS' ENTERED AT 09:55:57 ON 11 MAR 2003 SEL PN APPS L91

67 S E8+NT AND L122

E E3+ALL

E INTERLEUKINS/CT

L125

```
FILE 'WPIX' ENTERED AT 09:56:31 ON 11 MAR 2003
L93
              1 S E28-E35
L94
            135 S L4/BIX OR L7/BIX OR L8/BIX
                E FUCOID
L95
            138 S E3-E6, E8-E12/BIX
L96
            138 S L94, L95
                E RAOXPW/DCN
                E RAOXPW/DCN
             21 S E3-E11
L97
            138 S L96, L97
L98
              7 S L98 AND (A61P037 OR A61P043)/IC, ICM, ICS, ICA, ICI
L99
L100
              8 S L98 AND (B14-G02 OR B14-G02A OR C14-G02 OR B14-G02A OR B12-D
              5 S L98 AND (B14-L03 OR C14-L03 OR B14-L06 OR C14-L06 OR B12-G01
L101
L102
             13 S L98 AND D03-H01T?/MC
L103
              7 S L98 AND (P431 OR P617)/M0,M1,M2,M3,M4,M5,M6
L104
             19 S L98 AND (TAKARA? OR SHUZO?)/PA
L105
             16 S L98 AND (TOMINAGA ? OR YAMASHITA ? OR MIZUTANI ? OR SAGAWA ?
L106
              7 S L99-L103 AND L104,L105
L107
             19 S L99-L103 NOT L106
L108
             26 S L106, L107
L109
             14 S L104, L105 NOT L108
     FILE 'WPIX' ENTERED AT 10:08:09 ON 11 MAR 2003
     FILE 'FROSTI' ENTERED AT 10:08:52 ON 11 MAR 2003
L110
             16 S L4 OR L7 OR L8
                E FUCOI
             16 S E4, E8-E10
L111
L112
             16 S L110, L111
L113
              8 S L112 AND (ALLERG? OR FUNCTIONAL FOOD?)
L114
             11 S L112 AND PATENT
L115
             11 S L113, L114
L116
              5 S L112 NOT L115
     FILE 'FROSTI' ENTERED AT 10:11:20 ON 11 MAR 2003
     FILE 'FSTA' ENTERED AT 10:11:31 ON 11 MAR 2003
L117
             32 S L110
                E FUCOI
             32 S E4, E7, E8 OR L117
L118
     FILE 'MEDLINE' ENTERED AT 10:12:46 ON 11 MAR 2003
L119
            612 S L9
                E FUCOID
            513 S E4, E5, E8, E9, E12
L120
L121
            191 S E13-E15
L122
            619 S L119-L121
                E ALLERGY/CT
                E E3+ALL
                E E2+ALL
L123
              4 S E3+NT AND L122
L124
              5 S L10, L12, L13 AND L122
                E CYTOKINE/CT
```

Page		

## fonda 0 10 / 049419

L126		E25+NT AND L122
	E	INTERFERONS/CT
	Ε	E3+ALL
L127	5 S	E51+NT AND L122
	E	IGE/CT
	E	E3+ALL
	E	E2+ALL
L128	1 S	E46+NT AND L122
L129	0 S	E55+NT AND L122
L130	76 · S	L123-L128
L131	64 S	L130 AND PY<=2000
L132	2 S	L131 NOT AB/FA
L133	62 S	L131 NOT L132
	S	EL DN AN 7 27 32
L134	3 S	L133 AND E1-E9

FILE 'MEDLINE' ENTERED AT 10:20:53 ON 11 MAR 2003